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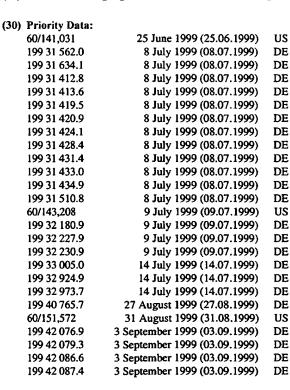
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(54) Title: CORYNEBACTERIUM GLUTAMICUM GENES ENCODING PROTEINS INVOLVED IN CARBON METABOLISM AND ENERGY PRODUCTION

(57) Abstract: Isolated nucleic acid molecules, designated SMP nucleic acid molecules, which encode novel SMP proteins from Corynebacterium glutamicum are described. The invention also provides antisense nucleic acid molecules, recombinant expression vectors containing SMP nucleic acid molecules, and host cells into which the expression vectors have been introduced. The invention still further provides isolated SMP proteins, mutated SMP proteins, fusion proteins, antigenic peptides and methods for the improvement of production of a desired compound from C. glutamicum based on genetic engineering of SMP genes in this organism.

# CORYNEBACTERIUM GLUTAMICUM GENES ENCODING PROTEINS INVOLVED IN CARBON METABOLISM AND ENERGY PRODUCTION

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#### **Related Applications**

5 This application claims priority to prior U.S. Provisional Patent Application Serial No. 60/141031, filed June 25, 1999, U.S. Provisional Patent Application Serial No. 60/143208, filed July 9, 1999, and U.S. Provisional Patent Application Serial No. 60/151572, filed August 31, 1999. This application also claims priority to prior German Patent Application No. 19931412.8, filed July 8, 1999, German Patent Application No. 19931413.6, filed July 8, 1999, German Patent Application No. 19931419.5, filed July 8, 1999, German Patent Application No. 19931420.9, filed July 8, 1999, German Patent Application No. 19931424.1, filed July 8, 1999, German Patent Application No. 19931428.4, filed July 8, 1999, German Patent Application No. 19931431.4, filed July 8, 1999, German Patent Application No. 19931433.0, filed July 8, 1999, German Patent Application No. 19931434.9, filed July 8, 1999, German Patent Application No. 19931510.8, filed July 8, 1999, German Patent Application No. 19931562.0, filed July 8, 1999, German Patent Application No. 19931634.1, filed July 8, 1999, German Patent Application No. 19932180.9, filed July 9, 1999, German Patent Application No. 19932227.9, filed July 9, 1999, German Patent Application No. 19932230.9, filed July 9, 1999, German Patent Application No. 19932924.9, filed July 14, 1999, German Patent Application No. 19932973.7, filed July 14, 1999, German Patent Application No. 19933005.0, filed July 14, 1999, German Patent Application No. 19940765.7, filed August 27, 1999, German Patent Application No. 19942076.9, filed September 3, 1999, German Patent Application No. 19942079.3, filed September 3, 1999, German Patent Application No. 19942086.6, filed September 3, 1999, German Patent Application No. 19942087.4, filed September 3, 1999, German Patent Application No. 19942088.2, filed September 3, 1999, German Patent Application No. 19942095.5, filed September 3, 1999, German Patent Application No. 19942123.4, filed September 3, 1999, and German Patent Application No. 19942125.0, filed September 3, 1999. The entire contents of all of the aforementioned application are hereby expressly incorporated 30 herein by this reference.

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#### **Background of the Invention**

Certain products and by-products of naturally-occurring metabolic processes in cells have utility in a wide array of industries, including the food, feed, cosmetics, and pharmaceutical industries. These molecules, collectively termed 'fine chemicals', include organic acids, both proteinogenic and non-proteinogenic amino acids, nucleotides and nucleosides, lipids and fatty acids, diols, carbohydrates, aromatic compounds, vitamins and cofactors, and enzymes. Their production is most conveniently performed through the large-scale culture of bacteria developed to produce and secrete large quantities of one or more desired molecules. One particularly useful organism for this purpose is *Corynebacterium glutamicum*, a gram positive, nonpathogenic bacterium. Through strain selection, a number of mutant strains have been developed which produce an array of desirable compounds. However, selection of strains improved for the production of a particular molecule is a time-consuming and difficult process.

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# Summary of the Invention

The invention provides novel bacterial nucleic acid molecules which have a variety of uses. These uses include the identification of microorganisms which can be used to produce fine chemicals, the modulation of fine chemical production in C. glutamicum or related bacteria, the typing or identification of C. glutamicum or related bacteria, as reference points for mapping the C. glutamicum genome, and as markers for transformation. These novel nucleic acid molecules encode proteins, referred to herein as sugar metabolism and oxidative phosphorylation (SMP) proteins.

C. glutamicum is a gram positive, aerobic bacterium which is commonly used in industry for the large-scale production of a variety of fine chemicals, and also for the degradation of hydrocarbons (such as in petroleum spills) and for the oxidation of terpenoids. The SMP nucleic acid molecules of the invention, therefore, can be used to identify microorganisms which can be used to produce fine chemicals, e.g., by fermentation processes. Modulation of the expression of the SMP nucleic acids of the invention, or modification of the sequence of the SMP nucleic acid molecules of the invention, can be used to modulate the production of one or more fine chemicals from a

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microorganism (e.g., to improve the yield or production of one or more fine chemicals from a Corynebacterium or Brevibacterium species).

The SMP nucleic acids of the invention may also be used to identify an organism as being Corynebacterium glutamicum or a close relative thereof, or to identify the presence of C. glutamicum or a relative thereof in a mixed population of microorganisms. The invention provides the nucleic acid sequences of a number of C. glutamicum genes; by probing the extracted genomic DNA of a culture of a unique or mixed population of microorganisms under stringent conditions with a probe spanning a region of a C. glutamicum gene which is unique to this organism, one can ascertain whether this organism is present. Although Corynebacterium glutamicum itself is nonpathogenic, it is related to species pathogenic in humans, such as Corynebacterium diphtheriae (the causative agent of diphtheria); the detection of such organisms is of significant clinical relevance.

The SMP nucleic acid molecules of the invention may also serve as reference points for mapping of the *C. glutamicum* genome, or of genomes of related organisms. Similarly, these molecules, or variants or portions thereof, may serve as markers for genetically engineered Corynebacterium or Brevibacterium species.

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The SMP proteins encoded by the novel nucleic acid molecules of the invention are capable of, for example, performing a function involved in the metabolism of carbon compounds such as sugars or in the generation of energy molecules by processes such as oxidative phosphorylation in *Corynebacterium glutamicum*. Given the availability of cloning vectors for use in *Corynebacterium glutamicum*, such as those disclosed in Sinskey *et al.*, U.S. Patent No. 4,649,119, and techniques for genetic manipulation of *C. glutamicum* and the related *Brevibacterium* species (*e.g.*, *lactofermentum*) (Yoshihama et al., *J. Bacteriol*. 162: 591-597 (1985); Katsumata *et al.*, *J. Bacteriol*. 159: 306-311 (1984); and Santamaria *et al.*, *J. Gen. Microbiol*. 130: 2237-2246 (1984)), the nucleic acid molecules of the invention may be utilized in the genetic engineering of this organism to make it a better or more efficient producer of one or more fine chemicals. This improved production or efficiency of production of a fine chemical may be due to a direct effect of manipulation of a gene of the invention, or it may be due to an indirect effect of such manipulation.

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There are a number of mechanisms by which the alteration of an SMP protein of the invention may directly affect the yield, production, and/or efficiency of production of a fine chemical from a C. glutamicum strain incorporating such an altered protein. The degradation of high-energy carbon molecules such as sugars, and the conversion of compounds such as NADH and FADH<sub>2</sub> to compounds containing high energy phosphate bonds via oxidative phosphorylation results in a number of compounds which themselves may be desirable fine chemicals, such as pyruvate, ATP, NADH, and a number of intermediate sugar compounds. Further, the energy molecules (such as ATP) and the reducing equivalents (such as NADH or NADPH) produced by these metabolic pathways are utilized in the cell to drive reactions which would otherwise be energetically unfavorable. Such unfavorable reactions include many biosynthetic pathways for fine chemicals. By improving the ability of the cell to utilize a particular sugar (e.g., by manipulating the genes encoding enzymes involved in the degradation and conversion of that sugar into energy for the cell), one may increase the amount of energy available to permit unfavorable, yet desired metabolic reactions (e.g., the biosynthesis of a desired fine chemical) to occur.

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The mutagenesis of one or more SMP genes of the invention may also result in SMP proteins having altered activities which indirectly impact the production of one or more desired fine chemicals from C. glutamicum. For example, by increasing the efficiency of utilization of one or more sugars (such that the conversion of the sugar to useful energy molecules is improved), or by increasing the efficiency of conversion of reducing equivalents to useful energy molecules (e.g., by improving the efficiency of oxidative phosphorylation, or the activity of the ATP synthase), one can increase the amount of these high-energy compounds available to the cell to drive normally unfavorable metabolic processes. These processes include the construction of cell walls, transcription, translation, and the biosynthesis of compounds necessary for growth and division of the cells (e.g., nucleotides, amino acids, vitamins, lipids, etc.) (Lengeler et al. (1999) Biology of Prokaryotes, Thieme Verlag: Stuttgart, p. 88-109; 913-918; 875-899). By improving the growth and multiplication of these engineered cells, it is possible to increase both the viability of the cells in large-scale culture, and also to improve their rate of division, such that a relatively larger number of cells can survive in fermentor culture. The yield, production, or efficiency of production may be increased, at least

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due to the presence of a greater number of viable cells, each producing the desired fine chemical. Also, many of the degradation products produced during sugar metabolism are utilized by the cell as precursors or intermediates in the production of other desirable products, such as fine chemicals. So, by increasing the ability of the cell to metabolize sugars, the number of these degradation products available to the cell for other processes should also be increased.

The invention provides novel nucleic acid molecules which encode proteins, referred to herein as SMP proteins, which are capable of, for example, performing a function involved in the metabolism of carbon compounds such as sugars and the generation of energy molecules by processes such as oxidative phosphorylation in *Corynebacterium glutamicum*. Nucleic acid molecules encoding an SMP protein are referred to herein as SMP nucleic acid molecules. In a preferred embodiment, the SMP protein participates in the conversion of carbon molecules and degradation products thereof to energy which is utilized by the cell for metabolic processes. Examples of such proteins include those encoded by the genes set forth in Table 1.

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Accordingly, one aspect of the invention pertains to isolated nucleic acid molecules (e.g., cDNAs, DNAs, or RNAs) comprising a nucleotide sequence encoding an SMP protein or biologically active portions thereof, as well as nucleic acid fragments suitable as primers or hybridization probes for the detection or amplification of SMPencoding nucleic acid (e.g., DNA or mRNA). In particularly preferred embodiments, the isolated nucleic acid molecule comprises one of the nucleotide sequences set forth as the odd-numbered SEQ ID NOs in the Sequence Listing (e.g., SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:5, SEQ ID NO:7....), or the coding region or a complement thereof of one of these nucleotide sequences. In other particularly preferred embodiments, the isolated nucleic acid molecule of the invention comprises a nucleotide sequence which hybridizes to or is at least about 50%, preferably at least about 60%, more preferably at least about 70%, 80% or 90%, and even more preferably at least about 95%, 96%, 97%, 98%, 99% or more homologous to a nucleotide sequence set forth as an odd-numbered SEQ ID NO in the Sequence Listing (e.g., SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:5, SEO ID NO:7....), or a portion thereof. In other preferred embodiments, the isolated nucleic acid molecule encodes one of the amino acid sequences set forth as an evennumbered SEQ ID NO in the Sequence Listing (e.g., SEQ ID NO:2, SEQ ID NO:4, SEQ

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ID NO:6, SEQ ID NO:8...).. The preferred SMP proteins of the present invention also preferably possess at least one of the SMP activities described herein.

In another embodiment, the isolated nucleic acid molecule encodes a protein or portion thereof wherein the protein or portion thereof includes an amino acid sequence which is sufficiently homologous to an amino acid sequence of the invention (e.g., a sequence having an even-numbered SEQ ID NO: in the Sequence Listing), e.g., sufficiently homologous to an amino acid sequence of the invention such that the protein or portion thereof maintains an SMP activity. Preferably, the protein or portion thereof encoded by the nucleic acid molecule maintains the ability to perform a function involved in the metabolism of carbon compounds such as sugars or the generation of energy molecules (e.g., ATP) by processes such as oxidative phosphorylation in Corynebacterium glutamicum. In one embodiment, the protein encoded by the nucleic acid molecule is at least about 50%, preferably at least about 60%, and more preferably at least about 70%, 80%, or 90% and most preferably at least about 95%, 96%, 97%, 98%, or 99% or more homologous to an amino acid sequence of the invention (e.g., an entire amino acid sequence selected those having an even-numbered SEQ ID NO in the Sequence Listing). In another preferred embodiment, the protein is a full length C. glutamicum protein which is substantially homologous to an entire amino acid sequence of the invention (encoded by an open reading frame shown in the corresponding oddnumbered SEQ ID NOs in the Sequence Listing (e.g., SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:5, SEQ ID NO:7....).

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In another preferred embodiment, the isolated nucleic acid molecule is derived from *C. glutamicum* and encodes a protein (*e.g.*, an SMP fusion protein) which includes a biologically active domain which is at least about 50% or more homologous to one of the amino acid sequences of the invention (*e.g.*, a sequence of one of the even-numbered SEQ ID NOs in the Sequence Listing) and is able to perform a function involved in the metabolism of carbon compounds such as sugars or the generation of energy molecules (*e.g.*, ATP) by processes such as oxidative phosphorylation in *Corynebacterium glutamicum*, or has one or more of the activities set forth in Table 1, and which also includes heterologous nucleic acid sequences encoding a heterologous polypeptide or regulatory regions.

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In another embodiment, the isolated nucleic acid molecule is at least 15 nucleotides in length and hybridizes under stringent conditions to a nucleic acid molecule comprising a nucleotide sequence of the invention (e.g., a sequence of an odd-numbered SEQ ID NO in the Sequence Listing) A. Preferably, the isolated nucleic acid molecule corresponds to a naturally-occurring nucleic acid molecule. More preferably, the isolated nucleic acid encodes a naturally-occurring C. glutamicum SMP protein, or a biologically active portion thereof.

Another aspect of the invention pertains to vectors, e.g., recombinant expression vectors, containing the nucleic acid molecules of the invention, and host cells into which such vectors have been introduced. In one embodiment, such a host cell is used to produce an SMP protein by culturing the host cell in a suitable medium. The SMP protein can be then isolated from the medium or the host cell.

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Yet another aspect of the invention pertains to a genetically altered microorganism in which an SMP gene has been introduced or altered. In one embodiment, the genome of the microorganism has been altered by introduction of a nucleic acid molecule of the invention encoding wild-type or mutated SMP sequence as a transgene. In another embodiment, an endogenous SMP gene within the genome of the microorganism has been altered, e.g., functionally disrupted, by homologous recombination with an altered SMP gene. In another embodiment, an endogenous or introduced SMP gene in a microorganism has been altered by one or more point mutations, deletions, or inversions, but still encodes a functional SMP protein. In still another embodiment, one or more of the regulatory regions (e.g., a promoter, repressor, or inducer) of an SMP gene in a microorganism has been altered (e.g., by deletion, truncation, inversion, or point mutation) such that the expression of the SMP gene is modulated. In a preferred embodiment, the microorganism belongs to the genus Corynebacterium or Brevibacterium, with Corynebacterium glutamicum being particularly preferred. In a preferred embodiment, the microorganism is also utilized for the production of a desired compound, such as an amino acid, with lysine being particularly preferred.

In another aspect, the invention provides a method of identifying the presence or activity of *Cornyebacterium diphtheriae* in a subject. This method includes detection of one or more of the nucleic acid or amino acid sequences of the invention (e.g., the

sequences set forth in the Sequence Listing as SEQ ID NOs 1 through 782) in a subject, thereby detecting the presence or activity of *Corynebacterium diphtheriae* in the subject.

Still another aspect of the invention pertains to an isolated SMP protein or a portion, e.g., a biologically active portion, thereof. In a preferred embodiment, the isolated SMP protein or portion thereof is capable of performing a function involved in the metabolism of carbon compounds such as sugars or in the generation of energy molecules (e.g., ATP) by processes such as oxidative phosphorylation in Corynebacterium glutamicum. In another preferred embodiment, the isolated SMP protein or portion thereof is sufficiently homologous to an amino acid sequence of the invention (e.g., a sequence of an even-numbered SEQ ID NO: in the Sequence Listing) such that the protein or portion thereof maintains the ability to perform a function involved in the metabolism of carbon compounds such as sugars or in the generation of energy molecules (e.g., ATP) by processes such as oxidative phosphorylation in Corynebacterium glutamicum.

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The invention also provides an isolated preparation of an SMP protein. In preferred embodiments, the SMP protein comprises an amino acid sequence of the invention (e.g., a sequence of an even-numbered SEQ ID NO: of the Sequence Listing). In another preferred embodiment, the invention pertains to an isolated full length protein which is substantially homologous to an entire amino acid sequence of the invention (e.g., a sequence of an even-numbered SEQ ID NO: of the Sequence Listing) (encoded by an open reading frame set forth in a corresponding odd-numbered SEQ ID NO: of the Sequence Listing). In yet another embodiment, the protein is at least about 50%, preferably at least about 60%, and more preferably at least about 70%, 80%, or 90%, and most preferably at least about 95%, 96%, 97%, 98%, or 99% or more homologous to an entire amino acid sequence of the invention (e.g., a sequence of an even-numbered SEQ ID NO: of the Sequence Listing). In other embodiments, the isolated SMP protein comprises an amino acid sequence which is at least about 50% or more homologous to one of the amino acid sequences of the invention (e.g., a sequence of an even-numbered SEQ ID NO: of the Sequence Listing) and is able to perform a function involved in the metabolism of carbon compounds such as sugars or in the generation of energy molecules (e.g., ATP) by processes such as oxidative phosphorylation in Corvnebacterium glutamicum, or has one or more of the activities set forth in Table 1.

Alternatively, the isolated SMP protein can comprise an amino acid sequence which is encoded by a nucleotide sequence which hybridizes, e.g., hybridizes under stringent conditions, or is at least about 50%, preferably at least about 60%, more preferably at least about 70%, 80%, or 90%, and even more preferably at least about 55%, 96%, 97%, 98,%, or 99% or more homologous to a nucleotide sequence of one of the even-numbered SEQ ID NOs set forth in the Sequence Listing. It is also preferred that the preferred forms of SMP proteins also have one or more of the SMP bioactivities described herein.

The SMP polypeptide, or a biologically active portion thereof, can be operatively linked to a non-SMP polypeptide to form a fusion protein. In preferred embodiments, this fusion protein has an activity which differs from that of the SMP protein alone. In other preferred embodiments, this fusion protein performs a function involved in the metabolism of carbon compounds such as sugars or in the generation of energy molecules (e.g., ATP) by processes such as oxidative phosphorylation in

15 Corynebacterium glutamicum. In particularly preferred embodiments, integration of this fusion protein into a host cell modulates production of a desired compound from the cell.

In another aspect, the invention provides methods for screening molecules which modulate the activity of an SMP protein, either by interacting with the protein itself or a substrate or binding partner of the SMP protein, or by modulating the transcription or translation of an SMP nucleic acid molecule of the invention.

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Another aspect of the invention pertains to a method for producing a fine chemical. This method involves the culturing of a cell containing a vector directing the expression of an SMP nucleic acid molecule of the invention, such that a fine chemical is produced. In a preferred embodiment, this method further includes the step of obtaining a cell containing such a vector, in which a cell is transfected with a vector directing the expression of an SMP nucleic acid. In another preferred embodiment, this method further includes the step of recovering the fine chemical from the culture. In a particularly preferred embodiment, the cell is from the genus *Corynebacterium* or *Brevibacterium*, or is selected from those strains set forth in Table 3.

Another aspect of the invention pertains to methods for modulating production of a molecule from a microorganism. Such methods include contacting the cell with an

agent which modulates SMP protein activity or SMP nucleic acid expression such that a cell associated activity is altered relative to this same activity in the absence of the agent. In a preferred embodiment, the cell is modulated for one or more *C. glutamicum* carbon metabolism pathways or for the production of energy through processes such as oxidative phosphorylation, such that the yields or rate of production of a desired fine chemical by this microorganism is improved. The agent which modulates SMP protein activity can be an agent which stimulates SMP protein activity or SMP nucleic acid expression. Examples of agents which stimulate SMP proteins, and nucleic acids encoding SMP proteins that have been introduced into the cell. Examples of agents which inhibit SMP activity or expression include small molecules and antisense SMP nucleic acid molecules.

Another aspect of the invention pertains to methods for modulating yields of a desired compound from a cell, involving the introduction of a wild-type or mutant SMP gene into a cell, either maintained on a separate plasmid or integrated into the genome of the host cell. If integrated into the genome, such integration can be random, or it can take place by homologous recombination such that the native gene is replaced by the introduced copy, causing the production of the desired compound from the cell to be modulated. In a preferred embodiment, said yields are increased. In another preferred embodiment, said chemical is a fine chemical. In a particularly preferred embodiment, said fine chemical is an amino acid. In especially preferred embodiments, said amino acid is L-lysine.

## **Detailed Description of the Invention**

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The present invention provides SMP nucleic acid and protein molecules which are involved in the metabolism of carbon compounds such as sugars and the generation of energy molecules by processes such as oxidative phosphorylation in Corynebacterium glutamicum. The molecules of the invention may be utilized in the modulation of production of fine chemicals from microorganisms, such as C.

30 glutamicum, either directly (e.g., where overexpression or optimization of a glycolytic pathway protein has a direct impact on the yield, production, and/or efficiency of production of, e.g., pyruvate from modified C. glutamicum), or may have an indirect

impact which nonetheless results in an increase of yield, production, and/or efficiency of production of the desired compound (e.g., where modulation of proteins involved in oxidative phosphorylation results in alterations in the amount of energy available to perform necessary metabolic processes and other cellular functions, such as nucleic acid and protein biosynthesis and transcription/translation). Aspects of the invention are further explicated below.

#### I. Fine Chemicals

The term 'fine chemical' is art-recognized and includes molecules produced by 10 an organism which have applications in various industries, such as, but not limited to, the pharmaceutical, agriculture, and cosmetics industries. Such compounds include organic acids, such as tartaric acid, itaconic acid, and diaminopimelic acid, both proteinogenic and non-proteinogenic amino acids, purine and pyrimidine bases, nucleosides, and nucleotides (as described e.g. in Kuninaka, A. (1996) Nucleotides and related compounds, p. 561-612, in Biotechnology vol. 6, Rehm et al., eds. VCH: 15 Weinheim, and references contained therein), lipids, both saturated and unsaturated fatty acids (e.g., arachidonic acid), diols (e.g., propane diol, and butane diol), carbohydrates (e.g., hyaluronic acid and trehalose), aromatic compounds (e.g., aromatic amines, vanillin, and indigo), vitamins and cofactors (as described in Ullmann's Encyclopedia of Industrial Chemistry, vol. A27, "Vitamins", p. 443-613 (1996) VCH: Weinheim and 20 references therein; and Ong, A.S., Niki, E. & Packer, L. (1995) "Nutrition, Lipids, Health, and Disease" Proceedings of the UNESCO/Confederation of Scientific and Technological Associations in Malaysia, and the Society for Free Radical Research -Asia, held Sept. 1-3, 1994 at Penang, Malaysia, AOCS Press, (1995)), enzymes, polyketides (Cane et al. (1998) Science 282: 63-68), and all other chemicals described in 25 Gutcho (1983) Chemicals by Fermentation, Noyes Data Corporation, ISBN: 0818805086 and references therein. The metabolism and uses of certain of these fine chemicals are further explicated below.

#### 30 A. Amino Acid Metabolism and Uses

Amino acids comprise the basic structural units of all proteins, and as such are essential for normal cellular functioning in all organisms. The term "amino acid" is art-

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recognized. The proteinogenic amino acids, of which there are 20 species, serve as structural units for proteins, in which they are linked by peptide bonds, while the nonproteinogenic amino acids (hundreds of which are known) are not normally found in proteins (see Ulmann's Encyclopedia of Industrial Chemistry, vol. A2, p. 57-97 VCH: Weinheim (1985)). Amino acids may be in the D- or L- optical configuration, though Lamino acids are generally the only type found in naturally-occurring proteins. Biosynthetic and degradative pathways of each of the 20 proteinogenic amino acids have been well characterized in both prokaryotic and eukaryotic cells (see, for example, Stryer, L. Biochemistry, 3<sup>rd</sup> edition, pages 578-590 (1988)). The 'essential' amino acids (histidine, isoleucine, leucine, lysine, methionine, phenylalanine, threonine, tryptophan, 10 and valine), so named because they are generally a nutritional requirement due to the complexity of their biosyntheses, are readily converted by simple biosynthetic pathways to the remaining 11 'nonessential' amino acids (alanine, arginine, asparagine, aspartate, cysteine, glutamate, glutamine, glycine, proline, serine, and tyrosine). Higher animals do retain the ability to synthesize some of these amino acids, but the essential amino 15 acids must be supplied from the diet in order for normal protein synthesis to occur.

Aside from their function in protein biosynthesis, these amino acids are interesting chemicals in their own right, and many have been found to have various applications in the food, feed, chemical, cosmetics, agriculture, and pharmaceutical industries. Lysine is an important amino acid in the nutrition not only of humans, but also of monogastric animals such as poultry and swine. Glutamate is most commonly used as a flavor additive (mono-sodium glutamate, MSG) and is widely used throughout the food industry, as are aspartate, phenylalanine, glycine, and cysteine. Glycine, L-methionine and tryptophan are all utilized in the pharmaceutical industry. Glutamine, valine, leucine, isoleucine, histidine, arginine, proline, serine and alanine are of use in both the pharmaceutical and cosmetics industries. Threonine, tryptophan, and D/L-methionine are common feed additives. (Leuchtenberger, W. (1996) Amino aids – technical production and use, p. 466-502 in Rehm *et al.* (eds.) Biotechnology vol. 6, chapter 14a, VCH: Weinheim). Additionally, these amino acids have been found to be useful as precursors for the synthesis of synthetic amino acids and proteins, such as N-acetylcysteine, S-carboxymethyl-L-cysteine, (S)-5-hydroxytryptophan, and others

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described in Ulmann's Encyclopedia of Industrial Chemistry, vol. A2, p. 57-97, VCH: Weinheim, 1985.

The biosynthesis of these natural amino acids in organisms capable of producing them, such as bacteria, has been well characterized (for review of bacterial amino acid biosynthesis and regulation thereof, see Umbarger, H.E.(1978) Ann. Rev. Biochem. 47: 533-606). Glutamate is synthesized by the reductive amination of αketoglutarate, an intermediate in the citric acid cycle. Glutamine, proline, and arginine are each subsequently produced from glutamate. The biosynthesis of serine is a threestep process beginning with 3-phosphoglycerate (an intermediate in glycolysis), and resulting in this amino acid after oxidation, transamination, and hydrolysis steps. Both cysteine and glycine are produced from serine; the former by the condensation of homocysteine with serine, and the latter by the transferal of the side-chain  $\beta$ -carbon atom to tetrahydrofolate, in a reaction catalyzed by serine transhydroxymethylase. Phenylalanine, and tyrosine are synthesized from the glycolytic and pentose phosphate pathway precursors erythrose 4-phosphate and phosphoenolpyruvate in a 9-step biosynthetic pathway that differ only at the final two steps after synthesis of prephenate. Tryptophan is also produced from these two initial molecules, but its synthesis is an 11step pathway. Tyrosine may also be synthesized from phenylalanine, in a reaction catalyzed by phenylalanine hydroxylase. Alanine, valine, and leucine are all biosynthetic products of pyruvate, the final product of glycolysis. Aspartate is formed from oxaloacetate, an intermediate of the citric acid cycle. Asparagine, methionine, threonine, and lysine are each produced by the conversion of aspartate. Isoleucine is formed from threonine. A complex 9-step pathway results in the production of histidine from 5-phosphoribosyl-1-pyrophosphate, an activated sugar.

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Amino acids in excess of the protein synthesis needs of the cell cannot be stored, and are instead degraded to provide intermediates for the major metabolic pathways of the cell (for review see Stryer, L. Biochemistry 3<sup>rd</sup> ed. Ch. 21 "Amino Acid Degradation and the Urea Cycle" p. 495-516 (1988)). Although the cell is able to convert unwanted amino acids into useful metabolic intermediates, amino acid production is costly in terms of energy, precursor molecules, and the enzymes necessary to synthesize them. Thus it is not surprising that amino acid biosynthesis is regulated by feedback inhibition, in which the presence of a particular amino acid serves to slow or entirely stop its own

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production (for overview of feedback mechanisms in amino acid biosynthetic pathways, see Stryer, L. Biochemistry, 3<sup>rd</sup> ed. Ch. 24: "Biosynthesis of Amino Acids and Heme" p. 575-600 (1988)). Thus, the output of any particular amino acid is limited by the amount of that amino acid present in the cell.

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## B. Vitamin, Cofactor, and Nutraceutical Metabolism and Uses

Vitamins, cofactors, and nutraceuticals comprise another group of molecules which the higher animals have lost the ability to synthesize and so must ingest, although they are readily synthesized by other organisms such as bacteria. These molecules are either bioactive substances themselves, or are precursors of biologically active substances which may serve as electron carriers or intermediates in a variety of metabolic pathways. Aside from their nutritive value, these compounds also have significant industrial value as coloring agents, antioxidants, and catalysts or other processing aids. (For an overview of the structure, activity, and industrial applications of these compounds, see, for example, Ullman's Encyclopedia of Industrial Chemistry, "Vitamins" vol. A27, p. 443-613, VCH: Weinheim, 1996.) The term "vitamin" is artrecognized, and includes nutrients which are required by an organism for normal functioning, but which that organism cannot synthesize by itself. The group of vitamins may encompass cofactors and nutraceutical compounds. The language "cofactor" includes nonproteinaceous compounds required for a normal enzymatic activity to occur. Such compounds may be organic or inorganic; the cofactor molecules of the invention are preferably organic. The term "nutraceutical" includes dietary supplements having health benefits in plants and animals, particularly humans. Examples of such molecules are vitamins, antioxidants, and also certain lipids (e.g., polyunsaturated fatty acids).

The biosynthesis of these molecules in organisms capable of producing them, such as bacteria, has been largely characterized (Ullman's Encyclopedia of Industrial Chemistry, "Vitamins" vol. A27, p. 443-613, VCH: Weinheim, 1996; Michal, G. (1999) Biochemical Pathways: An Atlas of Biochemistry and Molecular Biology, John Wiley & Sons; Ong, A.S., Niki, E. & Packer, L. (1995) "Nutrition, Lipids, Health, and Disease" Proceedings of the UNESCO/Confederation of Scientific and Technological

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Associations in Malaysia, and the Society for Free Radical Research – Asia, held Sept. 1-3, 1994 at Penang, Malaysia, AOCS Press: Champaign, IL X, 374 S).

Thiamin (vitamin B<sub>1</sub>) is produced by the chemical coupling of pyrimidine and thiazole mojeties. Riboflavin (vitamin B<sub>2</sub>) is synthesized from guanosine-5'-triphosphate (GTP) and ribose-5'-phosphate. Riboflavin, in turn, is utilized for the synthesis of flavin mononucleotide (FMN) and flavin adenine dinucleotide (FAD). The family of compounds collectively termed 'vitamin B6' (e.g., pyridoxine, pyridoxamine, pyridoxa-5'-phosphate, and the commercially used pyridoxin hydrochloride) are all derivatives of the common structural unit, 5-hydroxy-6-methylpyridine. Pantothenate (pantothenic acid, (R)-(+)-N-(2,4-dihydroxy-3,3-dimethyl-1-oxobutyl)-β-alanine) can be produced either by chemical synthesis or by fermentation. The final steps in pantothenate biosynthesis consist of the ATP-driven condensation of  $\beta$ -alanine and pantoic acid. The enzymes responsible for the biosynthesis steps for the conversion to pantoic acid, to βalanine and for the condensation to panthotenic acid are known. The metabolically active form of pantothenate is Coenzyme A, for which the biosynthesis proceeds in 5 enzymatic steps. Pantothenate, pyridoxal-5'-phosphate, cysteine and ATP are the precursors of Coenzyme A. These enzymes not only catalyze the formation of panthothante, but also the production of (R)-pantoic acid, (R)-pantolacton, (R)panthenol (provitamin B<sub>5</sub>), pantetheine (and its derivatives) and coenzyme A.

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Biotin biosynthesis from the precursor molecule pimeloyl-CoA in microorganisms has been studied in detail and several of the genes involved have been identified. Many of the corresponding proteins have been found to also be involved in Fe-cluster synthesis and are members of the nifS class of proteins. Lipoic acid is derived from octanoic acid, and serves as a coenzyme in energy metabolism, where it becomes part of the pyruvate dehydrogenase complex and the α-ketoglutarate dehydrogenase complex. The folates are a group of substances which are all derivatives of folic acid, which is turn is derived from L-glutamic acid, p-amino-benzoic acid and 6-methylpterin. The biosynthesis of folic acid and its derivatives, starting from the metabolism intermediates guanosine-5'-triphosphate (GTP), L-glutamic acid and p-amino-benzoic acid has been studied in detail in certain microorganisms.

Corrinoids (such as the cobalamines and particularly vitamin B<sub>12</sub>) and porphyrines belong to a group of chemicals characterized by a tetrapyrole ring system.

The biosynthesis of vitamin B<sub>12</sub> is sufficiently complex that it has not yet been completely characterized, but many of the enzymes and substrates involved are now known. Nicotinic acid (nicotinate), and nicotinamide are pyridine derivatives which are also termed 'niacin'. Niacin is the precursor of the important coenzymes NAD (nicotinamide adenine dinucleotide) and NADP (nicotinamide adenine dinucleotide phosphate) and their reduced forms.

The large-scale production of these compounds has largely relied on cell-free chemical syntheses, though some of these chemicals have also been produced by large-scale culture of microorganisms, such as riboflavin, Vitamin B<sub>6</sub>, pantothenate, and biotin. Only Vitamin B<sub>12</sub> is produced solely by fermentation, due to the complexity of its synthesis. *In vitro* methodologies require significant inputs of materials and time, often at great cost.

#### C. Purine, Pyrimidine, Nucleoside and Nucleotide Metabolism and Uses

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Purine and pyrimidine metabolism genes and their corresponding proteins are important targets for the therapy of tumor diseases and viral infections. The language "purine" or "pyrimidine" includes the nitrogenous bases which are constituents of nucleic acids, co-enzymes, and nucleotides. The term "nucleotide" includes the basic structural units of nucleic acid molecules, which are comprised of a nitrogenous base, a pentose sugar (in the case of RNA, the sugar is ribose; in the case of DNA, the sugar is D-deoxyribose), and phosphoric acid. The language "nucleoside" includes molecules which serve as precursors to nucleotides, but which are lacking the phosphoric acid moiety that nucleotides possess. By inhibiting the biosynthesis of these molecules, or their mobilization to form nucleic acid molecules, it is possible to inhibit RNA and DNA synthesis; by inhibiting this activity in a fashion targeted to cancerous cells, the ability of tumor cells to divide and replicate may be inhibited. Additionally, there are nucleotides which do not form nucleic acid molecules, but rather serve as energy stores (i.e., AMP) or as coenzymes (i.e., FAD and NAD).

Several publications have described the use of these chemicals for these medical indications, by influencing purine and/or pyrimidine metabolism (e.g. Christopherson, R.I. and Lyons, S.D. (1990) "Potent inhibitors of de novo pyrimidine and purine biosynthesis as chemotherapeutic agents." Med. Res. Reviews 10: 505-548). Studies of

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enzymes involved in purine and pyrimidine metabolism have been focused on the development of new drugs which can be used, for example, as immunosuppressants or anti-proliferants (Smith, J.L., (1995) "Enzymes in nucleotide synthesis." *Curr. Opin. Struct. Biol.* 5: 752-757; (1995) *Biochem Soc. Transact.* 23: 877-902). However, purine and pyrimidine bases, nucleosides and nucleotides have other utilities: as intermediates in the biosynthesis of several fine chemicals (e.g., thiamine, S-adenosyl-methionine, folates, or riboflavin), as energy carriers for the cell (e.g., ATP or GTP), and for chemicals themselves, commonly used as flavor enhancers (e.g., IMP or GMP) or for several medicinal applications (see, for example, Kuninaka, A. (1996) Nucleotides and Related Compounds in Biotechnology vol. 6, Rehm et al., eds. VCH: Weinheim, p. 561-612). Also, enzymes involved in purine, pyrimidine, nucleoside, or nucleotide metabolism are increasingly serving as targets against which chemicals for crop protection, including fungicides, herbicides and insecticides, are developed.

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The metabolism of these compounds in bacteria has been characterized (for reviews see, for example, Zalkin, H. and Dixon, J.E. (1992) "de novo purine nucleotide biosynthesis", in: Progress in Nucleic Acid Research and Molecular Biology, vol. 42, Academic Press:, p. 259-287; and Michal, G. (1999) "Nucleotides and Nucleosides", Chapter 8 in: Biochemical Pathways: An Atlas of Biochemistry and Molecular Biology, Wiley: New York). Purine metabolism has been the subject of intensive research, and is essential to the normal functioning of the cell. Impaired purine metabolism in higher animals can cause severe disease, such as gout. Purine nucleotides are synthesized from ribose-5-phosphate, in a series of steps through the intermediate compound inosine-5'phosphate (IMP), resulting in the production of guanosine-5'-monophosphate (GMP) or adenosine-5'-monophosphate (AMP), from which the triphosphate forms utilized as nucleotides are readily formed. These compounds are also utilized as energy stores, so their degradation provides energy for many different biochemical processes in the cell. Pyrimidine biosynthesis proceeds by the formation of uridine-5'-monophosphate (UMP) from ribose-5-phosphate. UMP, in turn, is converted to cytidine-5'-triphosphate (CTP). The deoxy- forms of all of these nucleotides are produced in a one step reduction reaction from the diphosphate ribose form of the nucleotide to the diphosphate deoxyribose form of the nucleotide. Upon phosphorylation, these molecules are able to participate in DNA synthesis.

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#### D. Trehalose Metabolism and Uses

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Trehalose consists of two glucose molecules, bound in  $\alpha$ ,  $\alpha$ -1,1 linkage. It is commonly used in the food industry as a sweetener, an additive for dried or frozen foods, and in beverages. However, it also has applications in the pharmaceutical, cosmetics and biotechnology industries (see, for example, Nishimoto et al., (1998) U.S. Patent No. 5,759,610; Singer, M.A. and Lindquist, S. (1998) Trends Biotech. 16: 460-467; Paiva, C.L.A. and Panek, A.D. (1996) Biotech. Ann. Rev. 2: 293-314; and Shiosaka, M. (1997) J. Japan 172: 97-102). Trehalose is produced by enzymes from many microorganisms and is naturally released into the surrounding medium, from which it can be collected using methods known in the art.

#### Sugar and Carbon Molecule Utilization and Oxidative Phosphorylation II.

Carbon is a critically important element for the formation of all organic compounds, and thus is a nutritional requirement not only for the growth and division of 15 C. glutamicum, but also for the overproduction of fine chemicals from this microorganism. Sugars, such as mono-, di-, or polysaccharides, are particularly good carbon sources, and thus standard growth media typically contain one or more of: glucose, fructose, mannose, galactose, ribose, sorbose, ribulose, lactose, maltose, sucrose, raffinose, starch, or cellulose (Ullmann's Encyclopedia of Industrial Chemistry 20 (1987) vol. A9, "Enzymes", VCH: Weinheim). Alternatively, more complex forms of sugar may be utilized in the media, such as molasses, or other by-products of sugar refinement. Other compounds aside from the sugars may be used as alternate carbon sources, including alcohols (e.g., ethanol or methanol), alkanes, sugar alcohols, fatty acids, and organic acids (e.g., acetic acid or lactic acid). For a review of carbon sources 25 and their utilization by microorganisms in culture, see: Ullman's Encyclopedia of Industrial Chemistry (1987) vol. A9, "Enzymes", VCH: Weinheim; Stoppok, E. and Buchholz, K. (1996) "Sugar-based raw materials for fermentation applications" in Biotechnology (Rehm, H.J. et al., eds.) vol. 6, VCH: Weinheim, p. 5-29; Rehm, H.J. (1980) Industrielle Mikrobiologie, Springer: Berlin; Bartholomew, W.H., and Reiman, 30 H.B. (1979). Economics of Fermentation Processes, in: Peppler, H.J. and Perlman, D., eds. Microbial Technology 2<sup>nd</sup> ed., vol. 2, chapter 18, Academic Press: New York; and

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Kockova-Kratachvilova, A. (1981) Characteristics of Industrial Microorganisms, in: Rehm, H.J. and Reed, G., eds. Handbook of Biotechnology, vol. 1, chapter 1, Verlag Chemie: Weinheim.

After uptake, these energy-rich carbon molecules must be processed such that

they are able to be degraded by one of the major sugar metabolic pathways. Such
pathways lead directly to useful degradation products, such as ribose-5-phosphate and
phosphoenolpyruvate, which may be subsequently converted to pyruvate. Three of the
most important pathways in bacteria for sugar metabolism include the EmbdenMeyerhoff-Pamas (EMP) pathway (also known as the glycolytic or fructose

bisphosphate pathway), the hexosemonophosphate (HMP) pathway (also known as the
pentose shunt or pentose phosphate pathway), and the Entner-Doudoroff (ED) pathway
(for review, see Michal, G. (1999) Biochemical Pathways: An Atlas of Biochemistry
and Molecular Biology, Wiley: New York, and Stryer, L. (1988) Biochemistry, Chapters
13-19, Freeman: New York, and references therein).

The EMP pathway converts hexose molecules to pyruvate, and in the process produces 2 molecules of ATP and 2 molecules of NADH. Starting with glucose-1-phosphate (which may be either directly taken up from the medium, or alternatively may be generated from glycogen, starch, or cellulose), the glucose molecule is isomerized to fructose-6-phosphate, is phosphorylated, and split into two 3-carbon molecules of glyceraldehyde-3-phosphate. After dehydrogenation, phosphorylation, and successive rearrangements, pyruvate results.

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The HMP pathway converts glucose to reducing equivalents, such as NADPH, and produces pentose and tetrose compounds which are necessary as intermediates and precursors in a number of other metabolic pathways. In the HMP pathway, glucose-6-phosphate is converted to ribulose-5-phosphate by two successive dehydrogenase reactions (which also release two NADPH molecules), and a carboxylation step. Ribulose-5-phosphate may also be converted to xyulose-5-phosphate and ribose-5-phosphate; the former can undergo a series of biochemical steps to glucose-6-phosphate, which may enter the EMP pathway, while the latter is commonly utilized as an intermediate in other biosynthetic pathways within the cell.

The ED pathway begins with the compound glucose or gluconate, which is subsequently phosphorylated and dehydrated to form 2-dehydro-3-deoxy-6-P-gluconate.

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Glucuronate and galacturonate may also be converted to 2-dehydro-3-deoxy-6-P-gluconate through more complex biochemical pathways. This product molecule is subsequently cleaved into glyceraldehyde-3-P and pyruvate; glyceraldehyde-3-P may itself also be converted to pyruvate.

The EMP and HMP pathways share many features, including intermediates and enzymes. The EMP pathway provides the greatest amount of ATP, but it does not produce ribose-5-phosphate, an important precursor for, e.g., nucleic acid biosynthesis, nor does it produce erythrose-4-phosphate, which is important for amino acid biosynthesis. Microorganisms that are capable of using only the EMP pathway for glucose utilization are thus not able to grow on simple media with glucose as the sole carbon source. They are referred to as fastidious organisms, and their growth requires inputs of complex organic compounds, such as those found in yeast extract.

In contrast, the HMP pathway produces all of the precursors necessary for both nucleic acid and amino acid biosynthesis, yet yields only half the amount of ATP energy that the EMP pathway does. The HMP pathway also produces NADPH, which may be used for redox reactions in biosynthetic pathways. The HMP pathway does not directly produce pyruvate, however, and thus these microorganisms must also possess this portion of the EMP pathway. It is therefore not surprising that a number of microorganisms, particularly the facultative anerobes, have evolved such that they possess both of these pathways.

The ED pathway has thus far has only been found in bacteria. Although this pathway is linked partly to the HMP pathway in the reverse direction for precursor formation, the ED pathway directly forms pyruvate by the aldolase cleavage of 3-ketodeoxy-6-phosphogluconate. The ED pathway can exist on its own and is utilized by the majority of strictly aerobic microorganisms. The net result is similar to that of the HMP pathway, although one mole of ATP can be formed only if the carbon atoms are converted into pyruvate, instead of into precursor molecules.

The pyruvate molecules produced through any of these pathways can be readily converted into energy via the Krebs cycle (also known as the citric acid cycle, the citrate cycle, or the tricarboxylic acid cycle (TCA cycle)). In this process, pyruvate is first decarboxylated, resulting in the production of one molecule of NADH, 1 molecule of acetyl-CoA, and 1 molecule of CO<sub>2</sub>. The acetyl group of acetyl CoA then reacts with

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the 4 carbon unit, oxaolacetate, leading to the formation of citric acid, a 6 carbon organic acid. Dehydration and two additional CO<sub>2</sub> molecules are released. Ultimately, oxaloacetate is regenerated and can serve again as an acetyl acceptor, thus completing the cycle. The electrons released during the oxidation of intermediates in the TCA cycle are transferred to NAD<sup>+</sup> to yield NADH.

During respiration, the electrons from NADH are transferred to molecular oxygen or other terminal electron acceptors. This process is catalyzed by the respiratory chain, an electron transport system containing both integral membrane proteins and membrane associated proteins. This system serves two basic functions: first, to accept electrons from an electron donor and to transfer them to an electron acceptor, and second, to conserve some of the energy released during electron transfer by the synthesis of ATP. Several types of oxidation-reduction enzymes and electron transport proteins are known to be involved in such processes, including the NADH dehydrogenases, flavin-containing electron carriers, iron sulfur proteins, and cytochromes. The NADH dehydrogenases are located at the cytoplasmic surface of the plasma membrane, and transfer hydrogen atoms from NADH to flavoproteins, in turn accepting electrons from NADH. The flavoproteins are a group of electron carriers possessing a flavin prosthetic group which is alternately reduced and oxidized as it accepts and transfers electrons. Three flavins are known to participate in these reactions: riboflavin, flavin-adenine dinucleotide (FAD) and flavin-mononucleotide (FMN). Iron sulfur proteins contain a cluster of iron and sulfur atoms which are not bonded to a heme group, but which still are able to participate in dehydration and rehydration reactions. Succinate dehydrogenase and aconitase are exemplary iron-sulfur proteins; their iron-sulfur complexes serve to accept and transfer electrons as part of the overall electron-transport chain. The cytochromes are proteins containing an iron porphyrin ring (heme). There are a number of different classes of cytochromes, differing in their reduction potentials. Functionally, these cytochromes form pathways in which electrons may be transferred to other cytochromes having increasingly more positive reduction potentials. A further class of non-protein electron carriers is known: the lipid-soluble quinones (e.g., 30 coenzyme Q). These molecules also serve as hydrogen atom acceptors and electron donors.

The action of the respiratory chain generates a proton gradient across the cell membrane, resulting in proton motive force. This force is utilized by the cell to synthesize ATP, via the membrane-spanning enzyme, ATP synthase. This enzyme is a multiprotein complex in which the transport of H<sup>+</sup> molecules through the membrane results in the physical rotation of the intracellular subunits and concomitant phosphorylation of ADP to form ATP (for review, see Fillingame, R.H. and Divall, S. (1999) *Novartis Found. Symp.* 221: 218-229, 229-234).

Non-hexose carbon substrates may also serve as carbon and energy sources for cells. Such substrates may first be converted to hexose sugars in the gluconeogenesis pathway, where glucose is first synthesized by the cell and then is degraded to produce energy. The starting material for this reaction is phosphoenolpyruvate (PEP), which is one of the key intermediates in the glycolytic pathway. PEP may be formed from substrates other than sugars, such as acetic acid, or by decarboxylation of oxaloacetate (itself an intermediate in the TCA cycle). By reversing the glycolytic pathway (utilizing a cascade of enzymes different than those of the original glycolysis pathway), glucose-6-phosphate may be formed. The conversion of pyruvate to glucose requires the utilization of 6 high energy phosphate bonds, whereas glycolysis only produces 2 ATP in the conversion of glucose to pyruvate. However, the complete oxidation of glucose (glycolysis, conversion of pyruvate into acetyl CoA, citric acid cycle, and oxidative phosphorylation) yields between 36-38 ATP, so the net loss of high energy phosphate bonds experienced during gluconeogenesis is offset by the overall greater gain in such high-energy molecules produced by the oxidation of glucose.

#### III. Elements and Methods of the Invention

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The present invention is based, at least in part, on the discovery of novel molecules, referred to herein as SMP nucleic acid and protein molecules, which participate in the conversion of sugars to useful degradation products and energy (e.g., ATP) in C. glutamicum or which may participate in the production of useful energy-rich molecules (e.g., ATP) by other processes, such as oxidative phosphorylation. In one embodiment, the SMP molecules participate in the metabolism of carbon compounds such as sugars or the generation of energy molecules (e.g., ATP) by processes such as oxidative phosphorylation in Corynebacterium glutamicum. In a preferred embodiment,

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the activity of the SMP molecules of the present invention to contribute to carbon metabolism or energy production in *C. glutamicum* has an impact on the production of a desired fine chemical by this organism. In a particularly preferred embodiment, the SMP molecules of the invention are modulated in activity, such that the *C. glutamicum* metabolic and energetic pathways in which the SMP proteins of the invention participate are modulated in yield, production, and/or efficiency of production, which either directly or indirectly modulates the yield, production, and/or efficiency of production of a desired fine chemical by *C. glutamicum*.

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The language, "SMP protein" or "SMP polypeptide" includes proteins which are capable of performing a function involved in the metabolism of carbon compounds such as sugars and the generation of energy molecules by processes such as oxidative phosphorylation in Corynebacterium glutamicum. Examples of SMP proteins include those encoded by the SMP genes set forth in Table 1 and by the odd-numbered SEO ID NOs. The terms "SMP gene" or "SMP nucleic acid sequence" include nucleic acid sequences encoding an SMP protein, which consist of a coding region and also corresponding untranslated 5' and 3' sequence regions. Examples of SMP genes include those set forth in Table 1. The terms "production" or "productivity" are art-recognized and include the concentration of the fermentation product (for example, the desired fine chemical) formed within a given time and a given fermentation volume (e.g., kg product per hour per liter). The term "efficiency of production" includes the time required for a. particular level of production to be achieved (for example, how long it takes for the cell to attain a particular rate of output of a fine chemical). The term "yield" or "product/carbon yield" is art-recognized and includes the efficiency of the conversion of the carbon source into the product (i.e., fine chemical). This is generally written as, for example, kg product per kg carbon source. By increasing the yield or production of the compound, the quantity of recovered molecules, or of useful recovered molecules of that compound in a given amount of culture over a given amount of time is increased. The terms "biosynthesis" or a "biosynthetic pathway" are art-recognized and include the synthesis of a compound, preferably an organic compound, by a cell from intermediate compounds in what may be a multistep and highly regulated process. The terms "degradation" or a "degradation pathway" are art-recognized and include the breakdown of a compound, preferably an organic compound, by a cell to degradation

products (generally speaking, smaller or less complex molecules) in what may be a multistep and highly regulated process. The term "degradation product" is art-recognized and includes breakdown products of a compound. Such products may themselves have utility as precursor (starting point) or intermediate molecules necessary for the biosynthesis of other compounds by the cell. The language "metabolism" is art-recognized and includes the totality of the biochemical reactions that take place in an organism. The metabolism of a particular compound, then, (e.g., the metabolism of an amino acid such as glycine) comprises the overall biosynthetic, modification, and degradation pathways in the cell related to this compound.

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In another embodiment, the SMP molecules of the invention are capable of modulating the production of a desired molecule, such as a fine chemical, in a microorganism such as C. glutamicum. There are a number of mechanisms by which the alteration of an SMP protein of the invention may directly affect the yield, production, and/or efficiency of production of a fine chemical from a C. glutamicum strain incorporating such an altered protein. The degradation of high-energy carbon molecules such as sugars, and the conversion of compounds such as NADH and FADH<sub>2</sub> to more useful forms via oxidative phosphorylation results in a number of compounds which themselves may be desirable fine chemicals, such as pyruvate, ATP, NADH, and a number of intermediate sugar compounds. Further, the energy molecules (such as ATP) and the reducing equivalents (such as NADH or NADPH) produced by these metabolic pathways are utilized in the cell to drive reactions which would otherwise be energetically unfavorable. Such unfavorable reactions include many biosynthetic pathways for fine chemicals. By improving the ability of the cell to utilize a particular sugar (e.g., by manipulating the genes encoding enzymes involved in the degradation and conversion of that sugar into energy for the cell), one may increase the amount of energy available to permit unfavorable, yet desired metabolic reactions (e.g., the biosynthesis of a desired fine chemical) to occur.

The mutagenesis of one or more SMP genes of the invention may also result in SMP proteins having altered activities which indirectly impact the production of one or more desired fine chemicals from *C. glutamicum*. For example, by increasing the efficiency of utilization of one or more sugars (such that the conversion of the sugar to useful energy molecules is improved), or by increasing the efficiency of conversion of

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reducing equivalents to useful energy molecules (e.g., by improving the efficiency of oxidative phosphorylation, or the activity of the ATP synthase), one can increase the amount of these high-energy compounds available to the cell to drive normally unfavorable metabolic processes. These processes include the construction of cell walls, transcription, translation, and the biosynthesis of compounds necessary for growth and division of the cells (e.g., nucleotides, amino acids, vitamins, lipids, etc.) (Lengeler et al. (1999) Biology of Prokaryotes, Thieme Verlag: Stuttgart, p. 88-109; 913-918; 875-899). By improving the growth and multiplication of these engineered cells, it is possible to increase both the viability of the cells in large-scale culture, and also to improve their rate of division, such that a relatively larger number of cells can survive in fermentor culture. The yield, production, or efficiency of production may be increased, at least due to the presence of a greater number of viable cells, each producing the desired fine chemical. Further, a number of the degradation and intermediate compounds produced during sugar metabolism are necessary precursors and intermediates for other biosynthetic pathways throughout the cell. For example, many amino acids are synthesized directly from compounds normally resulting from glycolysis or the TCA cycle (e.g., serine is synthesized from 3-phosphoglycerate, an intermediate in glycolysis). Thus, by increasing the efficiency of conversion of sugars to useful energy molecules, it is also possible to increase the amount of useful degradation products as well.

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The isolated nucleic acid sequences of the invention are contained within the genome of a Corynebacterium glutamicum strain available through the American Type Culture Collection, given designation ATCC 13032. The nucleotide sequence of the isolated C. glutamicum SMP DNAs and the predicted amino acid sequences of the C. glutamicum SMP proteins are shown in the Sequence Listing as odd-numbered SEQ ID NOs and even-numbered SEQ ID NOs, respectively. Computational analyses were performed which classified and/or identified these nucleotide sequences as sequences which encode proteins having a function involved in the metabolism of carbon compounds such as sugars or in the generation of energy molecules by processes such as oxidative phosphorylation in Corynebacterium glutamicum.

The present invention also pertains to proteins which have an amino acid sequence which is substantially homologous to an amino acid sequence of the invention

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(e.g., the sequence of an even-numbered SEQ ID NO of the Sequence Listing). As used herein, a protein which has an amino acid sequence which is substantially homologous to a selected amino acid sequence is least about 50% homologous to the selected amino acid sequence, e.g., the entire selected amino acid sequence. A protein which has an amino acid sequence which is substantially homologous to a selected amino acid sequence can also be least about 50-60%, preferably at least about 60-70%, and more preferably at least about 70-80%, 80-90%, or 90-95%, and most preferably at least about 96%, 97%, 98%, 99% or more homologous to the selected amino acid sequence.

An SMP protein or a biologically active portion or fragment thereof of the invention can participate in the metabolism of carbon compounds such as sugars or in the generation of energy molecules (e.g., ATP) by processes such as oxidative phosphorylation in *Corynebacterium glutamicum*, or can have one or more of the activities set forth in Table 1.

Various aspects of the invention are described in further detail in the following subsections:

#### A. Isolated Nucleic Acid Molecules

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One aspect of the invention pertains to isolated nucleic acid molecules that encode SMP polypeptides or biologically active portions thereof, as well as nucleic acid fragments sufficient for use as hybridization probes or primers for the identification or amplification of SMP-encoding nucleic acid (e.g., SMP DNA). As used herein, the term "nucleic acid molecule" is intended to include DNA molecules (e.g., cDNA or genomic DNA) and RNA molecules (e.g., mRNA) and analogs of the DNA or RNA generated using nucleotide analogs. This term also encompasses untranslated sequence located at both the 3' and 5' ends of the coding region of the gene: at least about 100 nucleotides of sequence upstream from the 5' end of the coding region and at least about 20 nucleotides of sequence downstream from the 3'end of the coding region of the gene. The nucleic acid molecule can be single-stranded or double-stranded, but preferably is double-stranded DNA. An "isolated" nucleic acid molecule is one which is separated from other nucleic acid molecules which are present in the natural source of the nucleic acid. Preferably, an "isolated" nucleic acid is free of sequences which naturally flank the nucleic acid (i.e., sequences located at the 5' and 3' ends of the nucleic acid) in the

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genomic DNA of the organism from which the nucleic acid is derived. For example, in various embodiments, the isolated SMP nucleic acid molecule can contain less than about 5 kb, 4kb, 3kb, 2kb, 1 kb, 0.5 kb or 0.1 kb of nucleotide sequences which naturally flank the nucleic acid molecule in genomic DNA of the cell from which the nucleic acid is derived (e.g, a *C. glutamicum* cell). Moreover, an "isolated" nucleic acid molecule, such as a DNA molecule, can be substantially free of other cellular material, or culture medium when produced by recombinant techniques, or chemical precursors or other chemicals when chemically synthesized.

A nucleic acid molecule of the present invention, e.g., a nucleic acid molecule 10 having a nucleotide sequence of an odd-numbered SEQ ID NO of the Sequence Listing, or a portion thereof, can be isolated using standard molecular biology techniques and the sequence information provided herein. For example, a C. glutamicum SMP DNA can be isolated from a C. glutamicum library using all or portion of one of the odd-numbered SEQ ID NO sequences of the Sequence Listing as a hybridization probe and standard hybridization techniques (e.g., as described in Sambrook, J., Fritsh, E. F., and Maniatis, T. Molecular Cloning: A Laboratory Manual. 2nd, ed., Cold Spring Harbor Laboratory, Cold Spring Harbor Laboratory Press, Cold Spring Harbor, NY, 1989). Moreover, a nucleic acid molecule encompassing all or a portion of one of the nucleic acid sequences of the invention (e.g., an odd-numbered SEQ ID NO:) can be isolated by the polymerase chain reaction using oligonucleotide primers designed based upon this 20 sequence (e.g., a nucleic acid molecule encompassing all or a portion of one of the nucleic acid sequences of the invention (e.g., an odd-numbered SEQ ID NO of the Sequence Listing) can be isolated by the polymerase chain reaction using oligonucleotide primers designed based upon this same sequence). For example, mRNA can be isolated from normal endothelial cells (e.g., by the guanidinium-thiocyanate 25 extraction procedure of Chirgwin et al. (1979) Biochemistry 18: 5294-5299) and DNA can be prepared using reverse transcriptase (e.g., Moloney MLV reverse transcriptase, available from Gibco/BRL, Bethesda, MD; or AMV reverse transcriptase, available from Seikagaku America, Inc., St. Petersburg, FL). Synthetic oligonucleotide primers for polymerase chain reaction amplification can be designed based upon one of the 30 nucleotide sequences shown in the Sequence Listing. A nucleic acid of the invention can be amplified using cDNA or, alternatively, genomic DNA, as a template and

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appropriate oligonucleotide primers according to standard PCR amplification techniques. The nucleic acid so amplified can be cloned into an appropriate vector and characterized by DNA sequence analysis. Furthermore, oligonucleotides corresponding to an SMP nucleotide sequence can be prepared by standard synthetic techniques, *e.g.*, using an automated DNA synthesizer.

In a preferred embodiment, an isolated nucleic acid molecule of the invention comprises one of the nucleotide sequences shown in the Sequence Listing. The nucleic acid sequences of the invention, as set forth in the Sequence Listing, correspond to the *Corynebacterium glutamicum* SMP DNAs of the invention. This DNA comprises sequences encoding SMP proteins (*i.e.*, the "coding region", indicated in each odd-numbered SEQ ID NO: sequence in the Sequence Listing), as well as 5' untranslated sequences and 3' untranslated sequences, also indicated in each odd-numbered SEQ ID NO: in the Sequence Listing. Alternatively, the nucleic acid molecule can comprise only the coding region of any of the sequences in nucleic acid sequences of the Sequence Listing.

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For the purposes of this application, it will be understood that each of the nucleic acid and amino acid sequences set forth in the Sequence Listing has an identifying RXA, RXN, or RXS number having the designation "RXA," "RXN," or "RXS" followed by 5 digits (i.e., RXA01626, RXN00043, or RXS0735). Each of the nucleic acid sequences comprises up to three parts: a 5' upstream region, a coding region, and a downstream region. Each of these three regions is identified by the same RXA, RXN, or RXS designation to eliminate confusion. The recitation "one of the odd-numbered sequences of the Sequence Listing", then, refers to any of the nucleic acid sequences in the Sequence Listing, which may also be distinguished by their differing RXA, RXN, or RXS designations. The coding region of each of these sequences is translated into a corresponding amino acid sequence, which is also set forth in the Sequence Listing, as an even-numbered SEQ ID NO: immediately following the corresponding nucleic acid sequence. For example, the coding region for RXA02735 is set forth in SEQ ID NO:1, while the amino acid sequence which it encodes is set forth as SEQ ID NO:2. The sequences of the nucleic acid molecules of the invention are identified by the same 30 RXA, RXN, or RXS designations as the amino acid molecules which they encode, such that they can be readily correlated. For example, the amino acid sequence designated

RXA00042 is a translation of the coding region of the nucleotide sequence of nucleic acid molecule RXA00042, and the amino acid sequence designated RXN00043 is a translation of the coding region of the nucleotide sequence of nucleic acid molecule RXN00043. The correspondence between the RXA, RXN and RXS nucleotide and amino acid sequences of the invention and their assigned SEQ ID NOs is set forth in Table 1.

Several of the genes of the invention are "F-designated genes". An F-designated gene includes those genes set forth in Table 1 which have an 'F' in front of the RXAdesignation. For example, SEQ ID NO:11, designated, as indicated on Table 1, as "F RXA01312", is an F-designated gene, as are SEQ ID NOs: 29, 33, and 39 (designated on Table 1 as "F RXA02803", "F RXA02854", and "F RXA01365", respectively).

In one embodiment, the nucleic acid molecules of the present invention are not intended to include those compiled in Table 2. In the case of the dapD gene, a sequence for this gene was published in Wehrmann, A., et al. (1998) J. Bacteriol. 180(12): 3159-3165. However, the sequence obtained by the inventors of the present application is significantly longer than the published version. It is believed that the published version relied on an incorrect start codon, and thus represents only a fragment of the actual coding region.

In another preferred embodiment, an isolated nucleic acid molecule of the invention comprises a nucleic acid molecule which is a complement of one of the nucleotide sequences of the invention (e.g., a sequence of an odd-numbered SEQ ID NO: of the Sequence Listing), or a portion thereof. A nucleic acid molecule which is complementary to one of the nucleotide sequences of the invention is one which is sufficiently complementary to one of the nucleotide sequences shown in the Sequence Listing (e.g., the sequence of an odd-numbered SEQ ID NO:) such that it can hybridize to one of the nucleotide sequences of the invention, thereby forming a stable duplex.

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In still another preferred embodiment, an isolated nucleic acid molecule of the invention comprises a nucleotide sequence which is at least about 50%, 51%, 52%, 53%, 54%, 55%, 56%, 57%, 58%, 59%, or 60%, preferably at least about 61%, 62%, 63%, 64%, 65%, 66%, 67%, 68%, 69%, or 70%, more preferably at least about 71%, 72%, 73%, 74%, 75%, 76%, 77%, 78%, 79%, or 80%, 81%, 82%, 83%, 84%, 85%, 86%,

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87%, 88%, 89%, or 90%, or 91%, 92%, 93%, 94%, and even more preferably at least about 95%, 96%, 97%, 98%, 99% or more homologous to a nucleotide sequence of the invention (e.g., a sequence of an odd-numbered SEQ ID NO: of the Sequence Listing), or a portion thereof. Ranges and identity values intermediate to the above-recited ranges, (e.g., 70-90% identical or 80-95% identical) are also intended to be encompassed by the present invention. For example, ranges of identity values using a combination of any of the above values recited as upper and/or lower limits are intendedto be included. In an additional preferred embodiment, an isolated nucleic acid molecule of the invention comprises a nucleotide sequence which hybridizes, e.g., hybridizes under stringent conditions, to one of the nucleotide sequences of the 10 invention, or a portion thereof.

Moreover, the nucleic acid molecule of the invention can comprise only a portion of the coding region of the sequence of one of the odd-numbered SEQ ID NOs of the Sequence Listing, for example a fragment which can be used as a probe or primer 15 · or a fragment encoding a biologically active portion of an SMP protein. The nucleotide sequences determined from the cloning of the SMP genes from C. glutamicum allows for the generation of probes and primers designed for use in identifying and/or cloning SMP homologues in other cell types and organisms, as well as SMP homologues from other Corynebacteria or related species. The probe/primer typically comprises substantially purified oligonucleotide. The oligonucleotide typically comprises a region of nucleotide sequence that hybridizes under stringent conditions to at least about 12, preferably about 25, more preferably about 40, 50 or 75 consecutive nucleotides of a sense strand of one of the nucleotide sequences of the invention (e.g., a sequence of one of the odd-numbered SEQ ID NOs of the Sequence Listing), an anti-sense sequence of one of these sequences, or naturally occurring mutants thereof. Primers based on a nucleotide sequence of the invention can be used in PCR reactions to clone SMP homologues. Probes based on the SMP nucleotide sequences can be used to detect transcripts or genomic sequences encoding the same or homologous proteins. In preferred embodiments, the probe further comprises a label group attached thereto, e.g. the label group can be a radioisotope, a fluorescent compound, an enzyme, or an enzyme co-factor. Such probes can be used as a part of a diagnostic test kit for identifying cells which misexpress an SMP protein, such as by measuring a level of an SMP-encoding

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nucleic acid in a sample of cells, e.g., detecting SMP mRNA levels or determining whether a genomic SMP gene has been mutated or deleted.

In one embodiment, the nucleic acid molecule of the invention encodes a protein or portion thereof which includes an amino acid sequence which is sufficiently homologous to an amino acid sequence of the invention (e.g., a sequence of an evennumbered SEQ ID NO of the Sequence Listing) such that the protein or portion thereof maintains the ability to perform a function involved in the metabolism of carbon compounds such as sugars or in the generation of energy molecules (e.g., ATP) by processes such as oxidative phosphorylation in Corynebacterium glutamicum. As used 10 herein, the language "sufficiently homologous" refers to proteins or portions thereof which have amino acid sequences which include a minimum number of identical or equivalent (e.g., an amino acid residue which has a similar side chain as an amino acid residue in a sequence of one of the even-numbered SEQ ID NOs of the Sequence Listing) amino acid residues to an amino acid sequence of the invention such that the protein or portion thereof is able to perform a function involved in the metabolism of carbon compounds such as sugars or in the generation of energy molecules (e.g., ATP) by processes such as oxidative phosphorylation in Corynebacterium glutamicum. Protein members of such sugar metabolic pathways or energy producing systems, as described herein, may play a role in the production and secretion of one or more fine chemicals. Examples of such activities are also described herein. Thus, "the function of an SMP protein" contributes either directly or indirectly to the yield, production, and/or efficiency of production of one or more fine chemicals. Examples of SMP protein activities are set forth in Table 1.

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In another embodiment, the protein is at least about 50-60%, preferably at least about 60-70%, and more preferably at least about 70-80%, 80-90%, 90-95%, and most preferably at least about 96%, 97%, 98%, 99% or more homologous to an entire amino acid sequence of the invention(e.g., a sequence of an even-numbered SEQ ID NO: of the Sequence Listing).

Portions of proteins encoded by the SMP nucleic acid molecules of the invention are preferably biologically active portions of one of the SMP proteins. As used herein, the term "biologically active portion of an SMP protein" is intended to include a portion, e.g., a domain/motif, of an SMP protein that participates in the metabolism of carbon

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compounds such as sugars, or in energy-generating pathways in *C. glutamicum*, or has an activity as set forth in Table 1. To determine whether an SMP protein or a biologically active portion thereof can participate in the metabolism of carbon compounds or in the production of energy-rich molecules in *C. glutamicum*, an assay of enzymatic activity may be performed. Such assay methods are well known to those of ordinary skill in the art, as detailed in Example 8 of the Exemplification.

Additional nucleic acid fragments encoding biologically active portions of an SMP protein can be prepared by isolating a portion of one of the amino acid sequences of the invention (e.g., a sequence of an even-numbered SEQ ID NO: of the Sequence Listing), expressing the encoded portion of the SMP protein or peptide (e.g., by recombinant expression in vitro) and assessing the activity of the encoded portion of the SMP protein or peptide.

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The invention further encompasses nucleic acid molecules that differ from one of the nucleotide sequences of the invention (e.g., a sequence of an odd-numbered SEQ ID NO: of the Sequence Listing) (and portions thereof) due to degeneracy of the genetic code and thus encode the same SMP protein as that encoded by the nucleotide sequences of the invention. In another embodiment, an isolated nucleic acid molecule of the invention has a nucleotide sequence encoding a protein having an amino acid sequence shown in the Sequence Listing (e.g., an even-numbered SEQ ID NO:). In a still further embodiment, the nucleic acid molecule of the invention encodes a full length C. glutamicum protein which is substantially homologous to an amino acid of the invention (encoded by an open reading frame shown in an odd-numbered SEQ ID NO: of the Sequence Listing).

It will be understood by one of ordinary skill in the art that in one embodiment the sequences of the invention are not meant to include the sequences of the prior art, such as those Genbank sequences set forth in Tables 2 or 4 which were available prior to the present invention. In one embodiment, the invention includes nucleotide and amino acid sequences having a percent identity to a nucleotide or amino acid sequence of the invention which is greater than that of a sequence of the prior art (e.g., a Genbank sequence (or the protein encoded by such a sequence) set forth in Tables 2 or 4). For example, the invention includes a nucleotide sequence which is greater than and/or at least 58% identical to the nucleotide sequence designated RXA00014 (SEQ ID NO:41),

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a nucleotide sequence which is greater than and/or at least % identical to the nucleotide sequence designated RXA00195 (SEQ ID NO:399), and a nucleotide sequence which is greater than and/or at least 42% identical to the nucleotide sequence designated RXA00196 (SEQ ID NO:401). One of ordinary skill in the art would be able to calculate the lower threshold of percent identity for any given sequence of the invention by examining the GAP-calculated percent identity scores set forth in Table 4 for each of the three top hits for the given sequence, and by subtracting the highest GAP-calculated percent identity from 100 percent. One of ordinary skill in the art will also appreciate that nucleic acid and amino acid sequences having percent identities greater than the lower threshold so calculated (e.g., at least 50%, 51%, 52%, 53%, 54%, 55%, 56%, 57%, 58%, 59%, or 60%, preferably at least about 61%, 62%, 63%, 64%, 65%, 66%, 67%, 68%, 69%, or 70%, more preferably at least about 71%, 72%, 73%, 74%, 75%, 76%, 77%, 78%, 79%, or 80%, 81%, 82%, 83%, 84%, 85%, 86%, 87%, 88%, 89%, or 90%, or 91%, 92%, 93%, 94%, and even more preferably at least about 95%, 96%, 97%, 98%, 99% or more identical) are also encompassed by the invention.

In addition to the *C. glutamicum* SMP nucleotide sequences set forth in the Sequence Listing as odd-numbered SEQ ID NOs, it will be appreciated by those of ordinary skill in the art that DNA sequence polymorphisms that lead to changes in the amino acid sequences of SMP proteins may exist within a population (*e.g.*, the *C. glutamicum* population). Such genetic polymorphism in the SMP gene may exist among individuals within a population due to natural variation. As used herein, the terms "gene" and "recombinant gene" refer to nucleic acid molecules comprising an open reading frame encoding an SMP protein, preferably a *C. glutamicum* SMP protein. Such natural variations can typically result in 1-5% variance in the nucleotide sequence of the SMP gene. Any and all such nucleotide variations and resulting amino acid polymorphisms in SMP that are the result of natural variation and that do not alter the functional activity of SMP proteins are intended to be within the scope of the invention.

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Nucleic acid molecules corresponding to natural variants and non-C. glutamicum homologues of the C. glutamicum SMP DNA of the invention can be isolated based on their homology to the C. glutamicum SMP nucleic acid disclosed herein using the C. glutamicum DNA, or a portion thereof, as a hybridization probe according to standard hybridization techniques under stringent hybridization conditions. Accordingly, in

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another embodiment, an isolated nucleic acid molecule of the invention is at least 15 nucleotides in length and hybridizes under stringent conditions to the nucleic acid molecule comprising a nucleotide sequence of of an odd-numbered SEQ ID NO: of the Sequence Listing. In other embodiments, the nucleic acid is at least 30, 50, 100, 250 or more nucleotides in length. As used herein, the term "hybridizes under stringent conditions" is intended to describe conditions for hybridization and washing under which nucleotide sequences at least 60% homologous to each other typically remain hybridized to each other. Preferably, the conditions are such that sequences at least about 65%, more preferably at least about 70%, and even more preferably at least about 75% or more homologous to each other typically remain hybridized to each other. Such stringent conditions are known to those of ordinary skill in the art and can be found in Current Protocols in Molecular Biology, John Wiley & Sons, N.Y. (1989), 6.3.1-6.3.6. A preferred, non-limiting example of stringent hybridization conditions are hybridization in 6X sodium chloride/sodium citrate (SSC) at about 45°C, followed by one or more washes in 0.2 X SSC, 0.1% SDS at 50-65°C. Preferably, an isolated nucleic acid molecule of the invention that hybridizes under stringent conditions to a nucleotide sequence of the invention corresponds to a naturally-occurring nucleic acid molecule. As used herein, a "naturally-occurring" nucleic acid molecule refers to an RNA or DNA molecule having a nucleotide sequence that occurs in nature (e.g., encodes a natural protein). In one embodiment, the nucleic acid encodes a natural C. glutamicum SMP protein.

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In addition to naturally-occurring variants of the SMP sequence that may exist in the population, one of ordinary skill in the art will further appreciate that changes can be introduced by mutation into a nucleotide sequence of the invention, thereby leading to changes in the amino acid sequence of the encoded SMP protein, without altering the functional ability of the SMP protein. For example, nucleotide substitutions leading to amino acid substitutions at "non-essential" amino acid residues can be made in a nucleotide sequence of the invention. A "non-essential" amino acid residue is a residue that can be altered from the wild-type sequence of one of the SMP proteins (e.g., an 30 even-numbered SEO ID NO: of the Sequence Listing) without altering the activity of said SMP protein, whereas an "essential" amino acid residue is required for SMP protein activity. Other amino acid residues, however, (e.g., those that are not conserved or only

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semi-conserved in the domain having SMP activity) may not be essential for activity and thus are likely to be amenable to alteration without altering SMP activity.

Accordingly, another aspect of the invention pertains to nucleic acid molecules encoding SMP proteins that contain changes in amino acid residues that are not essential for SMP activity. Such SMP proteins differ in amino acid sequence from a sequence of an even-numbered SEQ ID NO: of the Sequence Listing yet retain at least one of the SMP activities described herein. In one embodiment, the isolated nucleic acid molecule comprises a nucleotide sequence encoding a protein, wherein the protein comprises an amino acid sequence at least about 50% homologous to an amino acid sequence of the invention and is capable of participate in the metabolism of carbon compounds such as sugars, or in the biosynthesis of high-energy compounds in C. glutamicum, or has one or more activities set forth in Table 1. Preferably, the protein encoded by the nucleic acid molecule is at least about 50-60% homologous to the amino acid sequence of one of the odd-numbered SEQ ID NOs of the Sequence Listing, more preferably at least about 60-70% homologous to one of these sequences, even more preferably at least about 70-80%, 80-90%, 90-95% homologous to one of these sequences, and most preferably at least about 96%, 97%, 98%, or 99% homologous to one of the amino acid sequences of the invention.

To determine the percent homology of two amino acid sequences (e.g., one of the amino acid sequences of the invention and a mutant form thereof) or of two nucleic acids, the sequences are aligned for optimal comparison purposes (e.g., gaps can be introduced in the sequence of one protein or nucleic acid for optimal alignment with the other protein or nucleic acid). The amino acid residues or nucleotides at corresponding amino acid positions or nucleotide positions are then compared. When a position in one sequence (e.g., one of the amino acid sequences the invention) is occupied by the same amino acid residue or nucleotide as the corresponding position in the other sequence (e.g., a mutant form of the amino acid sequence), then the molecules are homologous at that position (i.e., as used herein amino acid or nucleic acid "homology" is equivalent to amino acid or nucleic acid "identity"). The percent homology between the two sequences is a function of the number of identical positions shared by the sequences (i.e., % homology = # of identical positions/total # of positions x 100).

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An isolated nucleic acid molecule encoding an SMP protein homologous to a protein sequence of the invention (e.g., a sequence of an even-numbered SEQ ID NO: of the Sequence Listing) can be created by introducing one or more nucleotide substitutions, additions or deletions into a nucleotide sequence of the invention such that one or more amino acid substitutions, additions or deletions are introduced into the encoded protein. Mutations can be introduced into one of the nucleotide sequences of the invention by standard techniques, such as site-directed mutagenesis and PCRmediated mutagenesis. Preferably, conservative amino acid substitutions are made at one or more predicted non-essential amino acid residues. A "conservative amino acid substitution" is one in which the amino acid residue is replaced with an amino acid residue having a similar side chain. Families of amino acid residues having similar side chains have been defined in the art. These families include amino acids with basic side chains (e.g., lysine, arginine, histidine), acidic side chains (e.g., aspartic acid, glutamic acid), uncharged polar side chains (e.g., glycine, asparagine, glutamine, serine, threonine, tyrosine, cysteine), nonpolar side chains (e.g., alanine, valine, leucine, isoleucine, proline, phenylalanine, methionine, tryptophan), beta-branched side chains (e.g., threonine, valine, isoleucine) and aromatic side chains (e.g., tyrosine, phenylalanine, tryptophan, histidine). Thus, a predicted nonessential amino acid residue in an SMP protein is preferably replaced with another amino acid residue from the same side chain family. Alternatively, in another embodiment, mutations can be introduced randomly along all or part of an SMP coding sequence, such as by saturation mutagenesis, and the resultant mutants can be screened for an SMP activity described herein to identify mutants that retain SMP activity. Following mutagenesis of the nucleotide sequence of one of the odd-numbered SEQ ID NOs of the Sequence Listing, the encoded protein can be expressed recombinantly and the activity of the protein can be determined using, for example, assays described herein (see Example 8 of the Exemplification).

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In addition to the nucleic acid molecules encoding SMP proteins described above, another aspect of the invention pertains to isolated nucleic acid molecules which are antisense thereto. An "antisense" nucleic acid comprises a nucleotide sequence which is complementary to a "sense" nucleic acid encoding a protein, e.g., complementary to the coding strand of a double-stranded DNA molecule or

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complementary to an mRNA sequence. Accordingly, an antisense nucleic acid can hydrogen bond to a sense nucleic acid. The antisense nucleic acid can be complementary to an entire SMP coding strand, or to only a portion thereof. In one embodiment, an antisense nucleic acid molecule is antisense to a "coding region" of the coding strand of a nucleotide sequence encoding an SMP protein. The term "coding region" refers to the region of the nucleotide sequence comprising codons which are translated into amino acid residues (e.g., the entire coding region of NO. 3 (RXA01626) comprises nucleotides 1 to 345). In another embodiment, the antisense nucleic acid molecule is antisense to a "noncoding region" of the coding strand of a nucleotide sequence encoding SMP. The term "noncoding region" refers to 5' and 3' sequences which flank the coding region that are not translated into amino acids (i.e., also referred to as 5' and 3' untranslated regions).

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Given the coding strand sequences encoding SMP disclosed herein (e.g., the sequences set forth as odd-numbered SEQ ID NOs in the Sequence Listing), antisense nucleic acids of the invention can be designed according to the rules of Watson and Crick base pairing. The antisense nucleic acid molecule can be complementary to the entire coding region of SMP mRNA, but more preferably is an oligonucleotide which is antisense to only a portion of the coding or noncoding region of SMP mRNA. For example, the antisense oligonucleotide can be complementary to the region surrounding the translation start site of SMP mRNA. An antisense oligonucleotide can be, for example, about 5, 10, 15, 20, 25, 30, 35, 40, 45 or 50 nucleotides in length. An antisense nucleic acid of the invention can be constructed using chemical synthesis and enzymatic ligation reactions using procedures known in the art. For example, an antisense nucleic acid (e.g., an antisense oligonucleotide) can be chemically synthesized using naturally occurring nucleotides or variously modified nucleotides designed to increase the biological stability of the molecules or to increase the physical stability of the duplex formed between the antisense and sense nucleic acids, e.g., phosphorothioate derivatives and acridine substituted nucleotides can be used. Examples of modified nucleotides which can be used to generate the antisense nucleic acid include 5fluorouracil, 5-bromouracil, 5-chlorouracil, 5-iodouracil, hypoxanthine, xanthine, 4acetylcytosine, 5-(carboxyhydroxylmethyl) uracil, 5-carboxymethylaminomethyl-2thiouridine, 5-carboxymethylaminomethyluracil, dihydrouracil, beta-D-

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galactosylqueosine, inosine, N6-isopentenyladenine, 1-methylguanine, 1-methylinosine, 2,2-dimethylguanine, 2-methyladenine, 2-methylguanine, 3-methylcytosine, 5-methylcytosine, N6-adenine, 7-methylguanine, 5-methylaminomethyluracil, 5-methoxyaminomethyl-2-thiouracil, beta-D-mannosylqueosine, 5'-

methoxycarboxymethyluracil, 5-methoxyuracil, 2-methylthio-N6-isopentenyladenine, uracil-5-oxyacetic acid (v), wybutoxosine, pseudouracil, queosine, 2-thiocytosine, 5-methyl-2-thiouracil, 2-thiouracil, 4-thiouracil, 5-methyluracil, uracil-5-oxyacetic acid methylester, uracil-5-oxyacetic acid (v), 5-methyl-2-thiouracil, 3-(3-amino-3-N-2-carboxypropyl) uracil, (acp3)w, and 2,6-diaminopurine. Alternatively, the antisense nucleic acid can be produced biologically using an expression vector into which a nucleic acid has been subcloned in an antisense orientation (*i.e.*, RNA transcribed from the inserted nucleic acid will be of an antisense orientation to a target nucleic acid of interest, described further in the following subsection).

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The antisense nucleic acid molecules of the invention are typically administered to a cell or generated *in situ* such that they hybridize with or bind to cellular mRNA and/or genomic DNA encoding an SMP protein to thereby inhibit expression of the protein, *e.g.*, by inhibiting transcription and/or translation. The hybridization can be by conventional nucleotide complementarity to form a stable duplex, or, for example, in the case of an antisense nucleic acid molecule which binds to DNA duplexes, through specific interactions in the major groove of the double helix. The antisense molecule can be modified such that it specifically binds to a receptor or an antigen expressed on a selected cell surface, *e.g.*, by linking the antisense nucleic acid molecule to a peptide or an antibody which binds to a cell surface receptor or antigen. The antisense nucleic acid molecule can also be delivered to cells using the vectors described herein. To achieve sufficient intracellular concentrations of the antisense molecules, vector constructs in which the antisense nucleic acid molecule is placed under the control of a strong prokaryotic, viral, or eukaryotic promoter are preferred.

In yet another embodiment, the antisense nucleic acid molecule of the invention is an  $\alpha$ -anomeric nucleic acid molecule. An  $\alpha$ -anomeric nucleic acid molecule forms specific double-stranded hybrids with complementary RNA in which, contrary to the usual  $\beta$ -units, the strands run parallel to each other (Gaultier *et al.* (1987) *Nucleic Acids*. *Res.* 15:6625-6641). The antisense nucleic acid molecule can also comprise a 2'-o-

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methylribonucleotide (Inoue et al. (1987) Nucleic Acids Res. 15:6131-6148) or a chimeric RNA-DNA analogue (Inoue et al. (1987) FEBS Lett. 215:327-330).

In still another embodiment, an antisense nucleic acid of the invention is a ribozyme. Ribozymes are catalytic RNA molecules with ribonuclease activity which are capable of cleaving a single-stranded nucleic acid, such as an mRNA, to which they have a complementary region. Thus, ribozymes (e.g., hammerhead ribozymes (described in Haselhoff and Gerlach (1988) Nature 334:585-591)) can be used to catalytically cleave SMP mRNA transcripts to thereby inhibit translation of SMP mRNA. A ribozyme having specificity for an SMP-encoding nucleic acid can be designed based upon the nucleotide sequence of an SMP cDNA disclosed herein (i.e., SEQ ID NO. 3 (RXA01626)). For example, a derivative of a Tetrahymena L-19 IVS RNA can be constructed in which the nucleotide sequence of the active site is complementary to the nucleotide sequence to be cleaved in an SMP-encoding mRNA. See, e.g., Cech et al. U.S. Patent No. 4,987,071 and Cech et al. U.S. Patent No. 5,116,742. Alternatively, SMP mRNA can be used to select a catalytic RNA having a 15 specific ribonuclease activity from a pool of RNA molecules. See, e.g., Bartel, D. and Szostak, J.W. (1993) Science 261:1411-1418.

Alternatively, SMP gene expression can be inhibited by targeting nucleotide sequences complementary to the regulatory region of an SMP nucleotide sequence (e.g., an SMP promoter and/or enhancers) to form triple helical structures that prevent transcription of an SMP gene in target cells. See generally, Helene, C. (1991)

Anticancer Drug Des. 6(6):569-84; Helene, C. et al. (1992) Ann. N.Y. Acad. Sci. 660:27-36; and Maher, L.J. (1992) Bioassays 14(12):807-15.

### 25 B. Recombinant Expression Vectors and Host Cells

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Another aspect of the invention pertains to vectors, preferably expression vectors, containing a nucleic acid encoding an SMP protein (or a portion thereof). As used herein, the term "vector" refers to a nucleic acid molecule capable of transporting another nucleic acid to which it has been linked. One type of vector is a "plasmid", which refers to a circular double stranded DNA loop into which additional DNA segments can be ligated. Another type of vector is a viral vector, wherein additional DNA segments can be ligated into the viral genome. Certain vectors are capable of

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autonomous replication in a host cell into which they are introduced (e.g., bacterial vectors having a bacterial origin of replication and episomal mammalian vectors). Other vectors (e.g., non-episomal mammalian vectors) are integrated into the genome of a host cell upon introduction into the host cell, and thereby are replicated along with the host genome. Moreover, certain vectors are capable of directing the expression of genes to which they are operatively linked. Such vectors are referred to herein as "expression vectors". In general, expression vectors of utility in recombinant DNA techniques are often in the form of plasmids. In the present specification, "plasmid" and "vector" can be used interchangeably as the plasmid is the most commonly used form of vector. However, the invention is intended to include such other forms of expression vectors, such as viral vectors (e.g., replication defective retroviruses, adenoviruses and adeno-

However, the invention is intended to include such other forms of expression vectors, such as viral vectors (e.g., replication defective retroviruses, adenoviruses and adenoassociated viruses), which serve equivalent functions.

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The recombinant expression vectors of the invention comprise a nucleic acid of the invention in a form suitable for expression of the nucleic acid in a host cell, which means that the recombinant expression vectors include one or more regulatory sequences, selected on the basis of the host cells to be used for expression, which is operatively linked to the nucleic acid sequence to be expressed. Within a recombinant expression vector, "operably linked" is intended to mean that the nucleotide sequence of interest is linked to the regulatory sequence(s) in a manner which allows for expression of the nucleotide sequence (e.g., in an in vitro transcription/translation system or in a host cell when the vector is introduced into the host cell). The term "regulatory sequence" is intended to include promoters, enhancers and other expression control elements (e.g., polyadenylation signals). Such regulatory sequences are described, for example, in Goeddel; Gene Expression Technology: Methods in Enzymology 185, Academic Press, San Diego, CA (1990). Regulatory sequences include those which direct constitutive expression of a nucleotide sequence in many types of host cell and those which direct expression of the nucleotide sequence only in certain host cells. Preferred regulatory sequences are, for example, promoters such as cos-, tac-, trp-, tet-, trp-tet-, lpp-, lac-, lpp-lac-, lacI<sup>q</sup>-, T7-, T5-, T3-, gal-, trc-, ara-, SP6-, arny, SPO2, λ-P<sub>R</sub>or  $\lambda$  P<sub>L</sub>, which are used preferably in bacteria. Additional regulatory sequences are, for example, promoters from yeasts and fungi, such as ADC1, MFα, AC, P-60, CYC1, GAPDH, TEF, rp28, ADH, promoters from plants such as CaMV/35S, SSU, OCS, lib4,

usp, STLS1, B33, nos or ubiquitin- or phaseolin-promoters. It is also possible to use artificial promoters. It will be appreciated by those of ordinary skill in the art that the design of the expression vector can depend on such factors as the choice of the host cell to be transformed, the level of expression of protein desired, etc. The expression vectors of the invention can be introduced into host cells to thereby produce proteins or peptides, including fusion proteins or peptides, encoded by nucleic acids as described herein (e.g., SMP proteins, mutant forms of SMP proteins, fusion proteins, etc.).

The recombinant expression vectors of the invention can be designed for expression of SMP proteins in prokaryotic or eukaryotic cells. For example, SMP genes can be expressed in bacterial cells such as C. glutamicum, insect cells (using baculovirus expression vectors), yeast and other fungal cells (see Romanos, M.A. et al. (1992) "Foreign gene expression in yeast: a review", Yeast 8: 423-488; van den Hondel, C.A.M.J.J. et al. (1991) "Heterologous gene expression in filamentous fungi" in: More Gene Manipulations in Fungi, J.W. Bennet & L.L. Lasure, eds., p. 396-428: Academic Press: San Diego; and van den Hondel, C.A.M.J.J. & Punt, P.J. (1991) "Gene transfer systems and vector development for filamentous fungi, in: Applied Molecular Genetics of Fungi, Peberdy, J.F. et al., eds., p. 1-28, Cambridge University Press: Cambridge), algae and multicellular plant cells (see Schmidt, R. and Willmitzer, L. (1988) High efficiency Agrobacterium tumefaciens - mediated transformation of Arabidopsis thaliana leaf and cotyledon explants" Plant Cell Rep: 583-586), or mammalian cells. Suitable host cells are discussed further in Goeddel, Gene Expression Technology: Methods in Enzymology 185, Academic Press, San Diego, CA (1990). Alternatively, the recombinant expression vector can be transcribed and translated in vitro, for example using T7 promoter regulatory sequences and T7 polymerase.

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Expression of proteins in prokaryotes is most often carried out with vectors containing constitutive or inducible promoters directing the expression of either fusion or non-fusion proteins. Fusion vectors add a number of amino acids to a protein encoded therein, usually to the amino terminus of the recombinant protein but also to the C-terminus or fused within suitable regions in the proteins. Such fusion vectors typically serve three purposes: 1) to increase expression of recombinant protein; 2) to increase the solubility of the recombinant protein; and 3) to aid in the purification of the recombinant protein by acting as a ligand in affinity purification. Often, in fusion

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expression vectors, a proteolytic cleavage site is introduced at the junction of the fusion moiety and the recombinant protein to enable separation of the recombinant protein from the fusion moiety subsequent to purification of the fusion protein. Such enzymes, and their cognate recognition sequences, include Factor Xa, thrombin and enterokinase.

Typical fusion expression vectors include pGEX (Pharmacia Biotech Inc; Smith, D.B. and Johnson, K.S. (1988) *Gene* 67:31-40), pMAL (New England Biolabs, Beverly, MA) and pRIT5 (Pharmacia, Piscataway, NJ) which fuse glutathione S-transferase (GST), maltose E binding protein, or protein A, respectively, to the target recombinant protein. In one embodiment, the coding sequence of the SMP protein is cloned into a pGEX expression vector to create a vector encoding a fusion protein comprising, from the N-terminus to the C-terminus, GST-thrombin cleavage site-X protein. The fusion protein can be purified by affinity chromatography using glutathione-agarose resin. Recombinant SMP protein unfused to GST can be recovered by cleavage of the fusion protein with thrombin.

15 Examples of suitable inducible non-fusion E. coli expression vectors include pTrc (Amann et al., (1988) Gene 69:301-315), pLG338, pACYC184, pBR322, pUC18, pUC19, pKC30, pRep4, pHS1, pHS2, pPLc236, pMBL24, pLG200, pUR290, pIN-III113-B1, \(\lambda gt11\), pBdCl, and pET 11d (Studier et al., Gene Expression Technology: Methods in Enzymology 185, Academic Press, San Diego, California (1990) 60-89; and Pouwels et al., eds. (1985) Cloning Vectors. Elsevier: New York IBSN 0 444 904018). 20 Target gene expression from the pTrc vector relies on host RNA polymerase transcription from a hybrid trp-lac fusion promoter. Target gene expression from the pET 11d vector relies on transcription from a T7 gn10-lac fusion promoter mediated by a coexpressed viral RNA polymerase (T7 gnl). This viral polymerase is supplied by 25 host strains BL21(DE3) or HMS174(DE3) from a resident  $\lambda$  prophage harboring a T7 gn1 gene under the transcriptional control of the lacUV 5 promoter. For transformation of other varieties of bacteria, appropriate vectors may be selected. For example, the plasmids pIJ101, pIJ364, pIJ702 and pIJ361 are known to be useful in transforming Streptomyces, while plasmids pUB110, pC194, or pBD214 are suited for transformation 30 of Bacillus species. Several plasmids of use in the transfer of genetic information into Corynebacterium include pHM1519, pBL1, pSA77, or pAJ667 (Pouwels et al., eds. (1985) Cloning Vectors. Elsevier: New York IBSN 0 444 904018).

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One strategy to maximize recombinant protein expression is to express the protein in a host bacteria with an impaired capacity to proteolytically cleave the recombinant protein (Gottesman, S., Gene Expression Technology: Methods in Enzymology 185, Academic Press, San Diego, California (1990) 119-128). Another strategy is to alter the nucleic acid sequence of the nucleic acid to be inserted into an expression vector so that the individual codons for each amino acid are those preferentially utilized in the bacterium chosen for expression, such as C. glutamicum (Wada et al. (1992) Nucleic Acids Res. 20:2111-2118). Such alteration of nucleic acid sequences of the invention can be carried out by standard DNA synthesis techniques.

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In another embodiment, the SMP protein expression vector is a yeast expression vector. Examples of vectors for expression in yeast *S. cerevisiae* include pYepSec1 (Baldari, *et al.*, (1987) *Embo J.* 6:229-234), 2 μ, pAG-1, Yep6, Yep13, pEMBLYe23, pMFa (Kurjan and Herskowitz, (1982) *Cell* 30:933-943), pJRY88 (Schultz *et al.*, (1987) *Gene* 54:113-123), and pYES2 (Invitrogen Corporation, San Diego, CA). Vectors and methods for the construction of vectors appropriate for use in other fungi, such as the filamentous fungi, include those detailed in: van den Hondel, C.A.M.J.J. & Punt, P.J. (1991) "Gene transfer systems and vector development for filamentous fungi, in: Applied Molecular Genetics of Fungi, J.F. Peberdy, *et al.*, eds., p. 1-28, Cambridge University Press: Cambridge, and Pouwels *et al.*, eds. (1985) Cloning Vectors. Elsevier: New York (IBSN 0 444 904018).

Alternatively, the SMP proteins of the invention can be expressed in insect cells using baculovirus expression vectors. Baculovirus vectors available for expression of proteins in cultured insect cells (e.g., Sf 9 cells) include the pAc series (Smith et al. (1983) Mol. Cell Biol. 3:2156-2165) and the pVL series (Lucklow and Summers (1989) Virology 170:31-39).

In another embodiment, the SMP proteins of the invention may be expressed in unicellular plant cells (such as algae) or in plant cells from higher plants (e.g., the spermatophytes, such as crop plants). Examples of plant expression vectors include those detailed in: Becker, D., Kemper, E., Schell, J. and Masterson, R. (1992) "New plant binary vectors with selectable markers located proximal to the left border", *Plant Mol. Biol.* 20: 1195-1197; and Bevan, M.W. (1984) "Binary *Agrobacterium* vectors for plant transformation", *Nucl. Acid. Res.* 12: 8711-8721, and include pLGV23, pGHlac+,

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pBIN19, pAK2004, and pDH51 (Pouwels et al., eds. (1985) Cloning Vectors. Elsevier: New York IBSN 0 444 904018).

In yet another embodiment, a nucleic acid of the invention is expressed in mammalian cells using a mammalian expression vector. Examples of mammalian expression vectors include pCDM8 (Seed, B. (1987) *Nature* 329:840) and pMT2PC (Kaufman *et al.* (1987) *EMBO J.* 6:187-195). When used in mammalian cells, the expression vector's control functions are often provided by viral regulatory elements. For example, commonly used promoters are derived from polyoma, Adenovirus 2, cytomegalovirus and Simian Virus 40. For other suitable expression systems for both prokaryotic and eukaryotic cells see chapters 16 and 17 of Sambrook, J., Fritsh, E. F., and Maniatis, T. *Molecular Cloning: A Laboratory Manual. 2nd, ed., Cold Spring Harbor Laboratory*, Cold Spring Harbor Laboratory Press, Cold Spring Harbor, NY, 1989.

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In another embodiment, the recombinant mammalian expression vector is capable of directing expression of the nucleic acid preferentially in a particular cell type (e.g., tissue-specific regulatory elements are used to express the nucleic acid). Tissuespecific regulatory elements are known in the art. Non-limiting examples of suitable tissue-specific promoters include the albumin promoter (liver-specific; Pinkert et al. (1987) Genes Dev. 1:268-277), lymphoid-specific promoters (Calame and Eaton (1988) Adv. Immunol. 43:235-275), in particular promoters of T cell receptors (Winoto and 20 Baltimore (1989) EMBO J. 8:729-733) and immunoglobulins (Banerji et al. (1983) Cell 33:729-740; Oueen and Baltimore (1983) Cell 33:741-748), neuron-specific promoters (e.g., the neurofilament promoter; Byrne and Ruddle (1989) PNAS 86:5473-5477), pancreas-specific promoters (Edlund et al. (1985) Science 230:912-916), and mammary gland-specific promoters (e.g., milk whey promoter; U.S. Patent No. 4,873,316 and European Application Publication No. 264,166). Developmentally-regulated promoters are also encompassed, for example the murine hox promoters (Kessel and Gruss (1990) Science 249:374-379) and the  $\alpha$ -fetoprotein promoter (Campes and Tilghman (1989) Genes Dev. 3:537-546).

The invention further provides a recombinant expression vector comprising a DNA molecule of the invention cloned into the expression vector in an antisense orientation. That is, the DNA molecule is operatively linked to a regulatory sequence in

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a manner which allows for expression (by transcription of the DNA molecule) of an RNA molecule which is antisense to SMP mRNA. Regulatory sequences operatively linked to a nucleic acid cloned in the antisense orientation can be chosen which direct the continuous expression of the antisense RNA molecule in a variety of cell types, for instance viral promoters and/or enhancers, or regulatory sequences can be chosen which direct constitutive, tissue specific or cell type specific expression of antisense RNA. The antisense expression vector can be in the form of a recombinant plasmid, phagemid or attenuated virus in which antisense nucleic acids are produced under the control of a high efficiency regulatory region, the activity of which can be determined by the cell type into which the vector is introduced. For a discussion of the regulation of gene expression using antisense genes see Weintraub, H. et al., Antisense RNA as a molecular tool for genetic analysis, Reviews - Trends in Genetics, Vol. 1(1) 1986.

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Another aspect of the invention pertains to host cells into which a recombinant expression vector of the invention has been introduced. The terms "host cell" and "recombinant host cell" are used interchangeably herein. It is understood that such terms refer not only to the particular subject cell but to the progeny or potential progeny of such a cell. Because certain modifications may occur in succeeding generations due to either mutation or environmental influences, such progeny may not, in fact, be identical to the parent cell, but are still included within the scope of the term as used herein.

A host cell can be any prokaryotic or eukaryotic cell. For example, an SMP protein can be expressed in bacterial cells such as *C. glutamicum*, insect cells, yeast or mammalian cells (such as Chinese hamster ovary cells (CHO) or COS cells). Other suitable host cells are known to one of ordinary skill in the art. Microorganisms related to *Corynebacterium glutamicum* which may be conveniently used as host cells for the nucleic acid and protein molecules of the invention are set forth in Table 3.

Vector DNA can be introduced into prokaryotic or eukaryotic cells via conventional transformation or transfection techniques. As used herein, the terms "transformation" and "transfection", "conjugation" and "transduction" are intended to refer to a variety of art-recognized techniques for introducing foreign nucleic acid (e.g., linear DNA or RNA (e.g., a linearized vector or a gene construct alone without a vector) or nucleic acid in the form of a vector (e.g., a plasmid, phage, phasmid, phagemid,

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transposon or other DNA) into a host cell, including calcium phosphate or calcium chloride co-precipitation, DEAE-dextran-mediated transfection, lipofection, natural competence, chemical-mediated transfer, or electroporation. Suitable methods for transforming or transfecting host cells can be found in Sambrook, et al. (Molecular Cloning: A Laboratory Manual. 2nd, ed., Cold Spring Harbor Laboratory, Cold Spring Harbor Laboratory Press, Cold Spring Harbor, NY, 1989), and other laboratory manuals.

For stable transfection of mammalian cells, it is known that, depending upon the expression vector and transfection technique used, only a small fraction of cells may integrate the foreign DNA into their genome. In order to identify and select these integrants, a gene that encodes a selectable marker (e.g., resistance to antibiotics) is generally introduced into the host cells along with the gene of interest. Preferred selectable markers include those which confer resistance to drugs, such as G418, hygromycin and methotrexate. Nucleic acid encoding a selectable marker can be introduced into a host cell on the same vector as that encoding an SMP protein or can be introduced on a separate vector. Cells stably transfected with the introduced nucleic acid can be identified by, for example, drug selection (e.g., cells that have incorporated the selectable marker gene will survive, while the other cells die).

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To create a homologous recombinant microorganism, a vector is prepared which contains at least a portion of an SMP gene into which a deletion, addition or substitution has been introduced to thereby alter, e.g., functionally disrupt, the SMP gene. Preferably, this SMP gene is a Corynebacterium glutamicum SMP gene, but it can be a homologue from a related bacterium or even from a mammalian, yeast, or insect source. In a preferred embodiment, the vector is designed such that, upon homologous recombination, the endogenous SMP gene is functionally disrupted (i.e., no longer encodes a functional protein; also referred to as a "knock out" vector). Alternatively, the vector can be designed such that, upon homologous recombination, the endogenous SMP gene is mutated or otherwise altered but still encodes functional protein (e.g., the upstream regulatory region can be altered to thereby alter the expression of the endogenous SMP protein). In the homologous recombination vector, the altered portion 30 of the SMP gene is flanked at its 5' and 3' ends by additional nucleic acid of the SMP gene to allow for homologous recombination to occur between the exogenous SMP gene carried by the vector and an endogenous SMP gene in a microorganism. The additional

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flanking SMP nucleic acid is of sufficient length for successful homologous recombination with the endogenous gene. Typically, several kilobases of flanking DNA (both at the 5' and 3' ends) are included in the vector (see *e.g.*, Thomas, K.R., and Capecchi, M.R. (1987) Cell 51: 503 for a description of homologous recombination vectors). The vector is introduced into a microorganism (*e.g.*, by electroporation) and cells in which the introduced SMP gene has homologously recombined with the endogenous SMP gene are selected, using art-known techniques.

In another embodiment, recombinant microorganisms can be produced which contain selected systems which allow for regulated expression of the introduced gene. For example, inclusion of an SMP gene on a vector placing it under control of the lac operon permits expression of the SMP gene only in the presence of IPTG. Such regulatory systems are well known in the art.

In another embodiment, an endogenous SMP gene in a host cell is disrupted (e.g., by homologous recombination or other genetic means known in the art) such that expression of its protein product does not occur. In another embodiment, an endogenous or introduced SMP gene in a host cell has been altered by one or more point mutations, deletions, or inversions, but still encodes a functional SMP protein. In still another embodiment, one or more of the regulatory regions (e.g., a promoter, repressor, or inducer) of an SMP gene in a microorganism has been altered (e.g., by deletion, truncation, inversion, or point mutation) such that the expression of the SMP gene is modulated. One of ordinary skill in the art will appreciate that host cells containing more than one of the described SMP gene and protein modifications may be readily produced using the methods of the invention, and are meant to be included in the present invention.

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A host cell of the invention, such as a prokaryotic or eukaryotic host cell in culture, can be used to produce (*i.e.*, express) an SMP protein. Accordingly, the invention further provides methods for producing SMP proteins using the host cells of the invention. In one embodiment, the method comprises culturing the host cell of invention (into which a recombinant expression vector encoding an SMP protein has been introduced, or into which genome has been introduced a gene encoding a wild-type or altered SMP protein) in a suitable medium until SMP protein is produced. In another

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embodiment, the method further comprises isolating SMP proteins from the medium or the host cell.

### C. Isolated SMP Proteins

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Another aspect of the invention pertains to isolated SMP proteins, and biologically active portions thereof. An "isolated" or "purified" protein or biologically active portion thereof is substantially free of cellular material when produced by recombinant DNA techniques, or chemical precursors or other chemicals when chemically synthesized. The language "substantially free of cellular material" includes preparations of SMP protein in which the protein is separated from cellular components of the cells in which it is naturally or recombinantly produced. In one embodiment, the language "substantially free of cellular material" includes preparations of SMP protein having less than about 30% (by dry weight) of non-SMP protein (also referred to herein as a "contaminating protein"), more preferably less than about 20% of non-SMP protein, still more preferably less than about 10% of non-SMP protein, and most preferably less than about 5% non-SMP protein. When the SMP protein or biologically active portion thereof is recombinantly produced, it is also preferably substantially free of culture medium, i.e., culture medium represents less than about 20%, more preferably less than about 10%, and most preferably less than about 5% of the volume of the protein preparation. The language "substantially free of chemical precursors or other chemicals" includes preparations of SMP protein in which the protein is separated from chemical precursors or other chemicals which are involved in the synthesis of the protein. In one embodiment, the language "substantially free of chemical precursors or other chemicals" includes preparations of SMP protein having less than about 30% (by dry weight) of chemical precursors or non-SMP chemicals, more preferably less than about 20% chemical precursors or non-SMP chemicals, still more preferably less than about 10% chemical precursors or non-SMP chemicals, and most preferably less than about 5% chemical precursors or non-SMP chemicals. In preferred embodiments, isolated proteins or biologically active portions thereof lack contaminating proteins from the same organism from which the SMP protein is derived. Typically, such proteins are produced by recombinant expression of, for example, a C. glutamicum SMP protein in a microorganism such as C. glutamicum.

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An isolated SMP protein or a portion thereof of the invention can participate in the metabolism of carbon compounds such as sugars, or in the production of energy compounds (e.g., by oxidative phosphorylation) utilized to drive unfavorable metabolic pathways, or has one or more of the activities set forth in Table 1. In preferred embodiments, the protein or portion thereof comprises an amino acid sequence which is sufficiently homologous to an amino acid sequence of the invention (e.g., a sequence of an even-numbered SEO ID NO: of the Sequence Listing) such that the protein or portion thereof maintains the ability to perform a function involved in the metabolism of carbon compounds such as sugars or in the generation of energy molecules by processes such as oxidative phosphorylation in Corynebacterium glutamicum. The portion of the protein is preferably a biologically active portion as described herein. In another preferred embodiment, an SMP protein of the invention has an amino acid sequence set forth as an even-numbered SEQ ID NO: of the Sequence Listing. In yet another preferred embodiment, the SMP protein has an amino acid sequence which is encoded by a nucleotide sequence which hybridizes, e.g., hybridizes under stringent conditions, to a nucleotide sequence of the invention (e.g., a sequence of an odd-numbered SEQ ID NO: of the Sequence Listing). In still another preferred embodiment, the SMP protein has an amino acid sequence which is encoded by a nucleotide sequence that is at least about 50%, 51%, 52%, 53%, 54%, 55%, 56%, 57%, 58%, 59%, or 60%, preferably at least about 61%, 62%, 63%, 64%, 65%, 66%, 67%, 68%, 69%, or 70%, more preferably at least about 71%, 72%, 73%, 74%, 75%, 76%, 77%, 78%, 79%, or 80%, 81%, 82%, 83%, 84%, 85%, 86%, 87%, 88%, 89%, or 90%, or 91%, 92%, 93%, 94%, and even more preferably at least about 95%, 96%, 97%, 98%, 99% or more homologous to one of the nucleic acid sequences of the invention, or a portion thereof. Ranges and identity values intermediate to the above-recited values, (e.g., 70-90% identical or 80-95% identical) are also intended to be encompassed by the present invention. For example, ranges of identity values using a combination of any of the above values recited as upper and/or lower limits are intended to be included. The preferred SMP proteins of the present invention also preferably possess at least one of the SMP activities described herein. For example, a preferred SMP protein of the present invention includes an amino acid sequence encoded by a nucleotide sequence which hybridizes, e.g., hybridizes under stringent conditions, to a nucleotide sequence of the invention, and

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which can perform a function involved in the metabolism of carbon compounds such as sugars or in the generation of energy molecules (e.g., ATP) by processes such as oxidative phosphorylation in *Corynebacterium glutamicum*, or which has one or more of the activities set forth in Table 1.

In other embodiments, the SMP protein is substantially homologous to an amino acid sequence of of the invention (e.g., a sequence of an even-numbered SEQ ID NO: of the Sequence Listing) and retains the functional activity of the protein of one of the amino acid sequences of the invention yet differs in amino acid sequence due to natural variation or mutagenesis, as described in detail in subsection I above. Accordingly, in another embodiment, the SMP protein is a protein which comprises an amino acid sequence which is at least about 50%, 51%, 52%, 53%, 54%, 55%, 56%, 57%, 58%, 59%, or 60%, preferably at least about 61%, 62%, 63%, 64%, 65%, 66%, 67%, 68%, 69%, or 70%, more preferably at least about 71%, 72%, 73%, 74%, 75%, 76%, 77%, 78%, 79%, or 80%, 81%, 82%, 83%, 84%, 85%, 86%, 87%, 88%, 89%, or 90%, or 91%, 92%, 93%, 94%, and even more preferably at least about 95%, 96%, 97%, 98%, 99% or more homologous to an entire amino acid sequence of the invention and which has at least one of the SMP activities described herein. Ranges and identity values intermediate to the above-recited values, (e.g., 70-90% identical or 80-95% identical) are also intended to be encompassed by the present invention. For example, ranges of identity values using a combination of any of the above values recited as upper and/or lower limits are intended to be included. In another embodiment, the invention pertains to a full length C. glutamicum protein which is substantially homologous to an entire amino acid sequence of the invention.

Biologically active portions of an SMP protein include peptides comprising

amino acid sequences derived from the amino acid sequence of an SMP protein, e.g., an
amino acid sequence of an even-numbered SEQ ID NO: of the Sequence Listing or the
amino acid sequence of a protein homologous to an SMP protein, which include fewer
amino acids than a full length SMP protein or the full length protein which is
homologous to an SMP protein, and exhibit at least one activity of an SMP protein.

Typically, biologically active portions (peptides, e.g., peptides which are, for example,
5, 10, 15, 20, 30, 35, 36, 37, 38, 39, 40, 50, 100 or more amino acids in length) comprise
a domain or motif with at least one activity of an SMP protein. Moreover, other

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biologically active portions, in which other regions of the protein are deleted, can be prepared by recombinant techniques and evaluated for one or more of the activities described herein. Preferably, the biologically active portions of an SMP protein include one or more selected domains/motifs or portions thereof having biological activity.

SMP proteins are preferably produced by recombinant DNA techniques. For example, a nucleic acid molecule encoding the protein is cloned into an expression vector (as described above), the expression vector is introduced into a host cell (as described above) and the SMP protein is expressed in the host cell. The SMP protein can then be isolated from the cells by an appropriate purification scheme using standard protein purification techniques. Alternative to recombinant expression, an SMP protein, polypeptide, or peptide can be synthesized chemically using standard peptide synthesis techniques. Moreover, native SMP protein can be isolated from cells (e.g., endothelial cells), for example using an anti-SMP antibody, which can be produced by standard techniques utilizing an SMP protein or fragment thereof of this invention.

The invention also provides SMP chimeric or fusion proteins. As used herein, an SMP "chimeric protein" or "fusion protein" comprises an SMP polypeptide operatively linked to a non-SMP polypeptide. An "SMP polypeptide" refers to a polypeptide having an amino acid sequence corresponding to an SMP protein, whereas a "non-SMP polypeptide" refers to a polypeptide having an amino acid sequence corresponding to a protein which is not substantially homologous to the SMP protein, e.g., a protein which is different from the SMP protein and which is derived from the same or a different organism. Within the fusion protein, the term "operatively linked" is intended to indicate that the SMP polypeptide and the non-SMP polypeptide are fused in-frame to each other. The non-SMP polypeptide can be fused to the N-terminus or C-terminus of the SMP polypeptide. For example, in one embodiment the fusion protein is a GST-SMP fusion protein in which the SMP sequences are fused to the C-terminus of the GST sequences. Such fusion proteins can facilitate the purification of recombinant SMP proteins. In another embodiment, the fusion protein is an SMP protein containing a heterologous signal sequence at its N-terminus. In certain host cells (e.g., mammalian host cells), expression and/or secretion of an SMP protein can be increased through use of a heterologous signal sequence.

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Preferably, an SMP chimeric or fusion protein of the invention is produced by standard recombinant DNA techniques. For example, DNA fragments coding for the different polypeptide sequences are ligated together in-frame in accordance with conventional techniques, for example by employing blunt-ended or stagger-ended termini for ligation, restriction enzyme digestion to provide for appropriate termini, filling-in of cohesive ends, as appropriate, alkaline phosphatase treatment to avoid undesirable joining, and enzymatic ligation. In another embodiment, the fusion gene can be synthesized by conventional techniques including automated DNA synthesizers. Alternatively, PCR amplification of gene fragments can be carried out using anchor primers which give rise to complementary overhangs between two consecutive gene fragments which can subsequently be annealed and reamplified to generate a chimeric gene sequence (see, for example, Current Protocols in Molecular Biology, Ausubel et al., eds. John Wiley & Sons: 1992). Moreover, many expression vectors are commercially available that already encode a fusion moiety (e.g., a GST polypeptide). An SMP-encoding nucleic acid can be cloned into such an expression vector such that the fusion moiety is linked in-frame to the SMP protein.

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Homologues of the SMP protein can be generated by mutagenesis, e.g., discrete point mutation or truncation of the SMP protein. As used herein, the term "homologue" refers to a variant form of the SMP protein which acts as an agonist or antagonist of the activity of the SMP protein. An agonist of the SMP protein can retain substantially the same, or a subset, of the biological activities of the SMP protein. An antagonist of the SMP protein can inhibit one or more of the activities of the naturally occurring form of the SMP protein, by, for example, competitively binding to a downstream or upstream member of the sugar molecule metabolic cascade or the energy-producing pathway which includes the SMP protein.

In an alternative embodiment, homologues of the SMP protein can be identified by screening combinatorial libraries of mutants, e.g., truncation mutants, of the SMP protein for SMP protein agonist or antagonist activity. In one embodiment, a variegated library of SMP variants is generated by combinatorial mutagenesis at the nucleic acid level and is encoded by a variegated gene library. A variegated library of SMP variants can be produced by, for example, enzymatically ligating a mixture of synthetic oligonucleotides into gene sequences such that a degenerate set of potential SMP

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sequences is expressible as individual polypeptides, or alternatively, as a set of larger fusion proteins (e.g., for phage display) containing the set of SMP sequences therein. There are a variety of methods which can be used to produce libraries of potential SMP homologues from a degenerate oligonucleotide sequence. Chemical synthesis of a degenerate gene sequence can be performed in an automatic DNA synthesizer, and the synthetic gene then ligated into an appropriate expression vector. Use of a degenerate set of genes allows for the provision, in one mixture, of all of the sequences encoding the desired set of potential SMP sequences. Methods for synthesizing degenerate oligonucleotides are known in the art (see, e.g., Narang, S.A. (1983) Tetrahedron 39:3; Itakura et al. (1984) Annu. Rev. Biochem. 53:323; Itakura et al. (1984) Science 198:1056; Ike et al. (1983) Nucleic Acid Res. 11:477.

In addition, libraries of fragments of the SMP protein coding can be used to generate a variegated population of SMP fragments for screening and subsequent selection of homologues of an SMP protein. In one embodiment, a library of coding sequence fragments can be generated by treating a double stranded PCR fragment of an SMP coding sequence with a nuclease under conditions wherein nicking occurs only about once per molecule, denaturing the double stranded DNA, renaturing the DNA to form double stranded DNA which can include sense/antisense pairs from different nicked products, removing single stranded portions from reformed duplexes by treatment with S1 nuclease, and ligating the resulting fragment library into an expression vector. By this method, an expression library can be derived which encodes N-terminal, C-terminal and internal fragments of various sizes of the SMP protein.

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Several techniques are known in the art for screening gene products of combinatorial libraries made by point mutations or truncation, and for screening cDNA libraries for gene products having a selected property. Such techniques are adaptable for rapid screening of the gene libraries generated by the combinatorial mutagenesis of SMP homologues. The most widely used techniques, which are amenable to high through-put analysis, for screening large gene libraries typically include cloning the gene library into replicable expression vectors, transforming appropriate cells with the resulting library of vectors, and expressing the combinatorial genes under conditions in which detection of a desired activity facilitates isolation of the vector encoding the gene whose product was detected. Recursive ensemble mutagenesis (REM), a new technique which enhances the

frequency of functional mutants in the libraries, can be used in combination with the screening assays to identify SMP homologues (Arkin and Yourvan (1992) *PNAS* 89:7811-7815; Delgrave *et al.* (1993) *Protein Engineering* 6(3):327-331).

In another embodiment, cell based assays can be exploited to analyze a variegated SMP library, using methods well known in the art.

### D. Uses and Methods of the Invention

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The nucleic acid molecules, proteins, protein homologues, fusion proteins, primers, vectors, and host cells described herein can be used in one or more of the following methods: identification of *C. glutamicum* and related organisms; mapping of genomes of organisms related to *C. glutamicum*; identification and localization of *C. glutamicum* sequences of interest; evolutionary studies; determination of SMP protein regions required for function; modulation of an SMP protein activity; modulation of the metabolism of one or more sugars; modulation of high-energy molecule production in a cell (*i.e.*, ATP, NADPH); and modulation of cellular production of a desired compound, such as a fine chemical.

The SMP nucleic acid molecules of the invention have a variety of uses. First, they may be used to identify an organism as being Corynebacterium glutamicum or a close relative thereof. Also, they may be used to identify the presence of C. glutamicum or a relative thereof in a mixed population of microorganisms. The invention provides the nucleic acid sequences of a number of C. glutamicum genes; by probing the extracted genomic DNA of a culture of a unique or mixed population of microorganisms under stringent conditions with a probe spanning a region of a C. glutamicum gene which is unique to this organism, one can ascertain whether this organism is present.

Although Corynebacterium glutamicum itself is nonpathogenic, it is related to

Although Corynebacterium glutamicum itself is nonpathogenic, it is related to pathogenic species, such as Corynebacterium diphtheriae. Corynebacterium diphtheriae is the causative agent of diphtheria, a rapidly developing, acute, febrile infection which involves both local and systemic pathology. In this disease, a local lesion develops in the upper respiratory tract and involves necrotic injury to epithelial cells; the bacilli secrete toxin which is disseminated through this lesion to distal susceptible tissues of the body. Degenerative changes brought about by the inhibition of protein synthesis in these tissues, which include heart, muscle, peripheral nerves, adrenals, kidneys, liver and

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spleen, result in the systemic pathology of the disease. Diphtheria continues to have high incidence in many parts of the world, including Africa, Asia, Eastern Europe and the independent states of the former Soviet Union. An ongoing epidemic of diphtheria in the latter two regions has resulted in at least 5,000 deaths since 1990.

In one embodiment, the invention provides a method of identifying the presence or activity of Cornyebacterium diphtheriae in a subject. This method includes detection of one or more of the nucleic acid or amino acid sequences of the invention (e.g., the sequences set forth as odd-numbered or even-numbered SEQ ID NOs, respectively, in the Sequence Listing) in a subject, thereby detecting the presence or activity of Corynebacterium diphtheriae in the subject. C. glutamicum and C. diphtheriae are related bacteria, and many of the nucleic acid and protein molecules in C. glutamicum are homologous to C. diphtheriae nucleic acid and protein molecules, and can therefore be used to detect C. diphtheriae in a subject.

The nucleic acid and protein molecules of the invention may also serve as markers for specific regions of the genome. This has utility not only in the mapping of the genome, but also for functional studies of *C. glutamicum* proteins. For example, to identify the region of the genome to which a particular *C. glutamicum* DNA-binding protein binds, the *C. glutamicum* genome could be digested, and the fragments incubated with the DNA-binding protein. Those which bind the protein may be additionally probed with the nucleic acid molecules of the invention, preferably with readily detectable labels; binding of such a nucleic acid molecule to the genome fragment enables the localization of the fragment to the genome map of *C. glutamicum*, and, when performed multiple times with different enzymes, facilitates a rapid determination of the nucleic acid sequence to which the protein binds. Further, the nucleic acid molecules of the invention may be sufficiently homologous to the sequences of related species such that these nucleic acid molecules may serve as markers for the construction of a genomic map in related bacteria, such as *Brevibacterium lactofermentum*.

The SMP nucleic acid molecules of the invention are also useful for evolutionary and protein structural studies. The metabolic and energy-releasing processes in which the molecules of the invention participate are utilized by a wide variety of prokaryotic and eukaryotic cells; by comparing the sequences of the nucleic acid molecules of the present invention to those encoding similar enzymes from other organisms, the

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evolutionary relatedness of the organisms can be assessed. Similarly, such a comparison permits an assessment of which regions of the sequence are conserved and which are not, which may aid in determining those regions of the protein which are essential for the functioning of the enzyme. This type of determination is of value for protein engineering studies and may give an indication of what the protein can tolerate in terms of mutagenesis without losing function.

Manipulation of the SMP nucleic acid molecules of the invention may result in the production of SMP proteins having functional differences from the wild-type SMP proteins. These proteins may be improved in efficiency or activity, may be present in greater numbers in the cell than is usual, or may be decreased in efficiency or activity.

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The invention provides methods for screening molecules which modulate the activity of an SMP protein, either by interacting with the protein itself or a substrate or binding partner of the SMP protein, or by modulating the transcription or translation of an SMP nucleic acid molecule of the invention. In such methods, a microorganism expressing one or more SMP proteins of the invention is contacted with one or more test compounds, and the effect of each test compound on the activity or level of expression of the SMP protein is assessed.

There are a number of mechanisms by which the alteration of an SMP protein of the invention may directly affect the yield, production, and/or efficiency of production of a fine chemical from a *C. glutamicum* strain incorporating such an altered protein. The degradation of high-energy carbon molecules such as sugars, and the conversion of compounds such as NADH and FADH<sub>2</sub> to more useful forms via oxidative phosphorylation results in a number of compounds which themselves may be desirable fine chemicals, such as pyruvate, ATP, NADH, and a number of intermediate sugar compounds. Further, the energy molecules (such as ATP) and the reducing equivalents (such as NADH or NADPH) produced by these metabolic pathways are utilized in the cell to drive reactions which would otherwise be energetically unfavorable. Such unfavorable reactions include many biosynthetic pathways for fine chemicals. By improving the ability of the cell to utilize a particular sugar (e.g., by manipulating the genes encoding enzymes involved in the degradation and conversion of that sugar into energy for the cell), one may increase the amount of energy available to permit

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unfavorable, yet desired metabolic reactions (e.g., the biosynthesis of a desired fine chemical) to occur.

Further, modulation of one or more pathways involved in sugar utilization permits optimization of the conversion of the energy contained within the sugar molecule to the production of one or more desired fine chemicals. For example, by reducing the activity of enzymes involved in, for example, gluconeogenesis, more ATP is available to drive desired biochemical reactions (such as fine chemical biosyntheses) in the cell. Also, the overall production of energy molecules from sugars may be modulated to ensure that the cell maximizes its energy production from each sugar molecule. Inefficient sugar utilization can lead to excess CO<sub>2</sub> production and excess energy, which may result in futile metabolic cycles. By improving the metabolism of sugar molecules, the cell should be able to function more efficiently, with a need for fewer carbon molecules. This should result in an improved fine chemical product: sugar molecule ratio (improved carbon yield), and permits a decrease in the amount of sugars that must be added to the medium in large-scale fermentor culture of such engineered *C. glutamicum*.

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The mutagenesis of one or more SMP genes of the invention may also result in SMP proteins having altered activities which indirectly impact the production of one or more desired fine chemicals from C. glutamicum. For example, by increasing the efficiency of utilization of one or more sugars (such that the conversion of the sugar to useful energy molecules is improved), or by increasing the efficiency of conversion of reducing equivalents to useful energy molecules (e.g., by improving the efficiency of oxidative phosphorylation, or the activity of the ATP synthase), one can increase the amount of these high-energy compounds available to the cell to drive normally unfavorable metabolic processes. These processes include the construction of cell walls, transcription, translation, and the biosynthesis of compounds necessary for growth and division of the cells (e.g., nucleotides, amino acids, vitamins, lipids, etc.) (Lengeler et al. (1999) Biology of Prokaryotes, Thieme Verlag: Stuttgart, p. 88-109; 913-918; 875-899). By improving the growth and multiplication of these engineered cells, it is possible to increase both the viability of the cells in large-scale culture, and also to improve their rate of division, such that a relatively larger number of cells can survive in fermentor culture. The yield, production, or efficiency of production may be increased, at least

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due to the presence of a greater number of viable cells, each producing the desired fine chemical.

Further, many of the degradation products produced during sugar metabolism are themselves utilized by the cell as precursors or intermediates for the production of a number of other useful compounds, some of which are fine chemicals. For example, pyruvate is converted into the amino acid alanine, and ribose-5-phosphate is an integral part of, for example, nucleotide molecules. The amount and efficiency of sugar metabolism, then, has a profound effect on the availability of these degradation products in the cell. By increasing the ability of the cell to process sugars, either in terms of efficiency of existing pathways (e.g., by engineering enzymes involved in these pathways such that they are optimized in activity), or by increasing the availability of the enzymes involved in such pathways (e.g., by increasing the number of these enzymes present in the cell), it is possible to also increase the availability of these degradation products in the cell, which should in turn increase the production of many different other desirable compounds in the cell (e.g., fine chemicals).

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The aforementioned mutagenesis strategies for SMP proteins to result in increased yields of a fine chemical from *C. glutamicum* are not meant to be limiting; variations on these strategies will be readily apparent to one of ordinary skill in the art. Using such strategies, and incorporating the mechanisms disclosed herein, the nucleic acid and protein molecules of the invention may be utilized to generate *C. glutamicum* or related strains of bacteria expressing mutated SMP nucleic acid and protein molecules such that the yield, production, and/or efficiency of production of a desired compound is improved. This desired compound may be any product produced by *C. glutamicum*, which includes the final products of biosynthesis pathways and intermediates of naturally-occurring metabolic pathways, as well as molecules which do not naturally occur in the metabolism of *C. glutamicum*, but which are produced by a *C. glutamicum* strain of the invention.

This invention is further illustrated by the following examples which should not be construed as limiting. The contents of all references, patent applications, patents, published patent applications, Tables, and the sequence listing cited throughout this application are hereby incorporated by reference.

# **TABLE 1: GENES IN THE APPLICATION**

### HMP:

				Table	Table 1 (continued)	(penu
Nucleic Acid	Amino Acid	Identification Code	Contig.	NT Start	NT Stop	Function
SEQ ID NO	SEQ ID NO					
37	38	RXN01365	W0091	1476	103	PHOSPHOGLUCOMUTASE (EC 5.4.2.2) / PHOSPHOMANNOMUTASE
39	40	F RXA01365	GR00397	897	4	(EC 3.4.2.8) PHOSPHOGLUCOMUTASE (EC 5.4.2.2) / PHOSPHOMANNOMUTASE
;	ç		, , ,		;	(EC 5.4.2.8)
-4-	77	KXA00098	GK00014	6252	8144	GLUCOSE-6-PHOSPHATE ISOMERASE (GPI) (EC 5.3.1.9)
3 4	‡ 4	2XX01909	0,00000	15.40	260	GEOCOOR-9-17000777 F 1000MF740R A (GTI A) (EC 5.6.1.9)
<b>7</b>	Ç 6	DY 60240	GR00039	2204	2047	TOCOPTION OF TOTAL MICHAEL (FC 9.4.2.1)
÷ €	9 0	DV A00293	02/00/20	1464	1167	TOORTOOLICENATE MICHAEL (EC. 9.4.2.1)
D •	2 2	1000000 1000000	2000000	1431	040	PRODUCTION (CONTRACT MOTAGE (CO. 9.4.2.1)
53	7 P	RXA00206	GR00032	6171	5134	PHOSPHOGETCERALE MOLISOE (EC 3.4.2.1)
22.02	56	RXA01243	GR00359	2302	3261	1-PHOSPHOFRUCTOKINASE (FC 2.7 1.56)
57	28	RXA01882	GR00538	1165	2154	1-PHOSPHOFRUCTOKINASE (EC 2.7.1.56)
29	9	RXA01702	GR00479	1397	366	FRUCTOSE-BISPHOSPHATE ALDOLASE (EC 4.1.2.13)
19	62	RXA02258	GR00654	26451	27227	TRIOSEPHOSPHATE ISOMERASE (EC 5.3.1.1)
63	64	RXN01225	VV0064	6382	4943	GLYCERALDEHYDE 3-PHOSPHATE DEHYDRÓGENASE (EC 1.2.1.12)
65	99	F RXA01225	GR00354	5302	6741	GLYCERALDEHYDE-3-PHOSPHATE DEHYDROGENASE HOMOLOG
29	89	RXA02256	GR00654	23934	24935	GLYCERALDEHYDE 3-PHOSPHATE DEHYDROGENASE (EC 1.2.1.12)
69	20	RXA02257	GR00654	25155	26369	PHOSPHOGLYCERATE KINASE (EC 2.7.2.3)
71	72	RXA00235	GR00036	2365	1091	ENOLASE (EC 4.2.1.11)
73	74	RXA01093	GR00306	1552	122	PYRUVATE KINASE (EC 2.7.1.40)
75	92	RXN02675	8600/	72801	70945	PYRUVATE KINASE (EC 2.7.1.40)
77	78	F RXA02675	GR00754	7	364	PYRUVATE KINASE (EC 2.7.1.40)
79	8	F RXA02695	GR00755	2949	4370	PYRUVATE KINASE (EC 2.7.1.40)
81	82	EXA00682	GR00179	5299	3401	PHOSPHOENOLPYRUVATE SYNTHASE (EC 2.7.9.2)
83	8 6	KXA00683	GK00179	6440	5349	PHOSPHOENOLPYROVATE SYNTHASE (EC 2.7.9.2)
32	<b>9</b>	KXN00635	VV0135	22/08 80/27	2/602	PYKOVATE DEHYDROGENASE (CYTOCHROME) (EC 1.2.2.2)
ò 6	8 6	F PY A00636	GR007 00	8 «	200	PTROVATE DESTURBUEINAGE (CTTOCHROME) (ECT.2.2.2)
5	86	RXN03044	W0019	1391	222	
93	26	F RXA02852	GR00852	<u>}</u> ~	281	
95	96	F RXA00268	GR00041	125	955	Ü
97	86	RXN03086	VV0049	2243	2650	PYRUVATE DEHYDROGENASE E1 COMPONENT (EC 1.2.4.1)
66	9	F RXA02887	GR10022	411	4	PYRUVATE DEHYDROGENASE E1 COMPONENT (EC 1.2.4.1)
101	102	RXN03043	VV0019	<b>-</b>	1362	PYRUVATE DEHYDROGENASE E1 COMPONENT (EC 1.2.4.1)
103	104	F RXA02897	GR10039	1291	ις.	PYRUVATE DEHYDROGENASE E1 COMPONENT (EC 1.2.4.1)
105	106	RXN03083	<b>W0047</b>	88	1110	DIHYDROLIPOAMIDE DEHYDROGENASE (EC 1.8.1.4)
107	108	F RXA02853	GR10001	83	1495	DIHYDROLIPOAMIDE DEHYDROGENASE (EC 1.8.1.4)
109	110	RXA02259	GR00654	27401	30172	PHOSPHOENOLPYRUVATE CARBOXYLASE (EC 4.1.1.31)
11	112	RXN02326	<b>VV0047</b>	4500	5315	PYRUVATE CARBOXYLASE (EC 6.4.1.1)
113	114	F RXA02326	GR00668	5338	4523	PYRUVATE CARBOXYLASE
115	116	RXN02327	VV0047	3533	4492	PYRUVATE CARBOXYLASE (EC 6.4.1.1)
117	118	F RXA02327	GR00668	6305	5346	PYRUVATE CARBOXYLASE
119	220	KXN02328	VV004/	1842	3437	PYKUVATE CARBOXYLASE (EC 6.4.1.1)
121	122	P KXA02328 RXN01048	GK00668	12530	5401 41318	PYKUVALE CARBOXYLASE (EC 6.4.1.1) MALIC ENZYME (EC 1.1.1.30)
3	+3-	250000	>>>>	15000	2	MALIO ENZINIC (LO 1.1.1.39)

inued)	Function	MALIC ENZYME (EC 1.1.1.39)	MALIC ENZYME (EC 1.1.1.39)	L-LACTATE DEHYDROGENASE (EC 1.1.1.27)	D-LACTATE DEHYDROGENASE (CYTOCHROME) (EC 1.1.2.4)	D-LACTATE DEHYDROGENASE (CYTOCHROME) (EC 1.1.2.4)	L-LACTATE DEHYDROGENASE (CYTOCHROME) (EC 1.1.2.3)	D-LACTATE DEHYDROGENASE (EC 1.1.1.28)	D-LACTATE DEHYDROGENASE (EC 1.1.1.28)	D-LACTATE DEHYDROGENASE (EC 1.1.1.28)	D-3-PHOSPHOGLYCERATE DEHYDROGENASE (EC 1.1.1.95)	IOLB PROTEIN	IOLB PROTEIN: D-FRUCTOSE 1,6-BISPHOSPHATE = GLYCERONE-CC	PHOSPHATE + D- GLYCERALDEHYDE 3-PHOSPHATE.	IOLS PROTEIN	IOLS PROTEIN	NAGD PROTEIN	PUTATIVE N-GLYCERALDEHYDE-2-PHOSPHOTRANSFERASE	GLPX PROTEIN	D-3-PHOSPHOGLYCERATE DEHYDROGENASE (EC 1.1.1.95)	D-3-PHOSPHOGLYCERATE DEHYDROGENASE (EC 1.1.1.95)	PHOSPHOGLYCERATE MUTASE (EC 5.4.2.1)	PYRUVATE CARBOXYLASE (EC 6.4.1.1)	PYRUVATE DEHYDROGENASE E1 COMPONENT (EC 1.2.4.1)	PYRUVATE DEHYDROGENASE E1 COMPONENT (EC 1.2.4.1)	PHOSPHOENOLPYRUVATE CARBOXYKINASE [GTP] (EC 4.1.1.32)	LIPOAMIDE DEHYDROGENASE COMPONENT (E3) OF BRANCHED. CHAIN AI PHA-KETO ACID DEHYDROGENASE COMPLEX (EC. 1.8.1.4)	LIPOAMIDE DEHYDROGENASE COMPONENT (E3) OF BRANCHED- CHAIN ALPHA-KETO ACID DEHYDROGENASE COMPLEX (EC 1.8.1.4)				
Table 1 (continued)	NT Stop	290	5655	2820	38606	2837	5417	11666	216	6209	1734	5536	304	9	1116	2240	3207		559	295	8298	2074	2989	5224	686	58385	3428	519	281	12541	2296	3533
Table	NT Start	<del>ن</del> ق	4693	1879	35763	က	4158	9954	<del>-</del>	4611	2645	6138	7	509	568	3127	2344		287	287	7474	1250	3993	6135	1390	59053	3216	310	೮	14370	3477	3703
	Contig.	GR00296	GR00046	GR00755	VV0176	GR00048	GR00544	VV0105	GR00562	GR00562	GR00047	W0157	GR00315	VV0085	GR00316	VV0127	GR00239		VV0354	GR00816	VV0019	GR00422	GR00211	VV0213	GR00690	W0098	VV0052	VV0377	VV0382	VV0098	6000/\	6000/\
	Identification Code	F RXA01048	F RXA00290	RXA02694	RXN00296	F RXA00296	RXA01901	RXN01952	F RXA01952	F RXA01955	RXA00293	RXN01130	F RXA01130	RXN03112	F RXA01133	RXN00871	F RXA00871		RXN02829	F RXA02829	RXN01468	F RXA01468.	RXA00794	RXN02920	F RXA02379	RXN02688	RXN03087	RXN03186	RXN03187	RXN02591	RXS01260	RXS01261
	Amino Acid SEQ ID NO	126																	158	160	162	164	166	168	170	172	174	176	178	180	182	184
	Nucleic Acid SEQ ID NO	125	127	129	131	133	135	137	139	141	143	145	147	149	151	153	155		157	159	161	163	165	167	169	171	173	175	177	179	181	183

metabolism	
Glycerol 1	

Function		GLYCEROL KINASE (EC 2.7.1.30)	GLYCEROL-3-PHOSPHATE DEHYDROGENASE (NAD(P)+) (EC 1.1.1.94)	GLYCEROL-3-PHOSPHATE DEHYDROGENASE (NAD(P)+) (EC 1.1.1.94)	AEROBIC GLYCEROL-3-PHOSPHATE DEHYDROGENASE (EC 1.1.99.5)	GLYCEROL-3-PHOSPHATE REGULON REPRESSOR	GLYCEROL-3-PHOSPHATE REGULON REPRESSOR
NT Stop		2926	4488	1853	1830	2302	147
NT Start		1400	5483	939	3515	1526	392
Contig.		GR00749	W0143	GR00293	GR00525	GR00359	GR00661
Identification Code		RXA02640	RXN01025	F RXA01025	RXA01851	RXA01242	RXA02288
Amino Acid	SEQ ID NO						
Nucleic Acid	SEQ ID NO	185	187	189	191	193	195

(panul	Function		GLYCEROL-3-PHOSPHATE-BINDING PERIPLASMIC PROTEIN	PRECURSOR	GLYCEROL-3-PHOSPHATE-BINDING PERIPLASMIC PROTEIN	PRECURSOR	Uncharacterized protein involved in glycerol metabolism (homolog of	Drosophila rhomboid)	Glycerophosphoryl diester phosphodiesterase	
Table 1 (continued)	NT Start NT Stop Function		24086		918		3062		22807	
Table	NT Start		24949		1736		3808		22091	
	Contig.		W0122		GR00541		GR00703		VV0122	
	Identification Code		RXN01891		F RXA01891		RXA02414		RXN01580	
	Amino Acid	SEQ ID NO	198		200		202		204	
	Nucleic Acid	SEQ ID NO	197		199		201		203	

## Acetate metabolism

Function	ACETATE KINASE (EC 2.7.2.1)	ACETATE OPERON REPRESSOR	ALCOHOL DEHYDROGENASE (EC 1.1.1.1)	ALDEHYDE DEHYDROGENASE (EC	ALDEHYDE DEHYDROGENASE (EC 1.2.1.3)	ACETOLACTATE SYNTHASE LARGE SUBUNIT (EC 4.1.3.18)	ACETOLACTATE SYNTHASE LARGE SUBUNIT (EC 4,1,3,18)	ACETOLACTATE SYNTHASE LARGE SUBUNIT (EC 4.1.3.18)	ACETOLACTATE SYNTHASE SMALL SUBUNIT (EC 4.1.3.18)							
NT Stop	1357	7941	3391	1959	2419	2945	10159	437	10055	860	3160	14163	320	8254	935	7722
NT Start	2547	8744	4425	1360	1928	3961	11676	108	10678	က	1598	15614	2230	9372	243	8237
Contig.	GR00418	GR00179	GR00037	GR00438	GR00438	GR00498	GR00726	VV0034	W0155	VV0033	W0008	W0315	VV0127	7,0077	VV0264	7.000
Identification Code	RXA01436	RXA00686	RXA00246	RXA01571	RXA01572	RXA01758	RXA02539	RXN03061	RXN03150	RXN01340	RXN01498	RXN02674	RXN00868	RXN01143	RXN01146	RXN01144
Amino Acid SEQ ID NO	206	208	210	212	214	216	218	220	222	224	226	228	230	232	234	236
Nucleic Acid	205	207	209	211	213	215	217	219	221	223	225	227	229	231	233	235

# Butanediol, diacetyl and acetoin formation

Function	(S,S)-butane-2,3-diol dehydrogenase (EC 1.1	ACETOIN(DIACETYL) REDUCTASE (EC 1.1	ALCOHOL DEHYDROGENASE (EC 1.1.1)
NT Stop	7309	5351	28399
NT Start	8082	6103	27383
Contig.	GR00715	GR00710	W0112
Identification Code	RXA02474	RXA02453	RXS01758
Amino Acid SEQ ID NO	238	240	242
Nucleic Acid SEQ ID NO	237	239	241

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Function	GLUCOSE-6-PHOSPHATE 1-DEHYDROGENASE (EC 1.1.1.49)	TRANSALDOLASE (EC 2.2.1.2)	TRANSKETOLASE (EC 2.2.1.1)	6-PHOSPHOGLUCONATE DEHYDROGENASE, DECARBOXYLATING (EC	1.1.1.44) PHOSPHOGLUCONATE DEHYDROGENASE, DECARBOXYLATING (EC 4.4.4.4.4	6-PHOSPHOGLUCONATE DEHYDROGENASE, DECARBOXYLATING (EC 1.1.1.44)
NT Start NT Stop	1771	3420	. 4670	510	1366	4448
NT Start	3312	4499	69/9	1232	2817	3012
Contig.	_	_	_	GR00270	VV0106	GR00283
Identification Code	RXA02737	RXA02738	RXA02739	RXA00965	RXN00999	F RXA00999
Amino Acid SEQ ID NO						
Nucleic Acid	243	245	247	249	251	253

## Nucleotide sugar conversion

Function UDP-GAI ACTOPYRANOSE MITASE (FC 5 4 99 9)	UDP-GALACTOPYRANOSE MUTASE (EC 5.4.99.9)	UDP-GALACTOPYRANOSE MUTASE (EC 5.4.99.9)	UDP-GLUCOSE 6-DEHYDROGENASE (EC 1.1.1.22)	UDP-N-ACETYLENOLPYRUVOYLGLUCOSAMINE REDUCTASE (EC 1.1.1.158)	UDP-N-ACETYLGLUCOSAMINE PYROPHOSPHORYLASE (EC 2.7.7.23)	UTPGLUCOSE-1-PHOSPHATE URIDYLYLTRANSFERASE (EC 2.7.7.9)	UTP-GLUCOSE-1-PHOSPHATE URIDYLYLTRANSFERASE (EC 2.7.7.9)	GDP-MANNOSE 6-DEHYDROGENASE (EC 1.1.1.132)	MANNOSE-1-PHOSPHATE GUANYLTRANSFERASE (EC 2.7.7.13)	GLUCOSE-1-PHOSPHATE ADENYLYLTRANSFERASE (EC 2.7.7.27)	GLUCOSE-1-PHOSPHATE THYMIDYLYLTRANSFERASE (EC 2.7.7.24)	GLUCOSE-1-PHOSPHATE THYMIDYLYLTRANSFERASE (EC 2.7.7.24)	GLUCOSE-1-PHOSPHATE THYMIDYLYLTRANSFERASE (EC 2.7.7.24)	D-RIBITOL-5-PHOSPHATE CYTIDYLYLTRANSFERASE (EC 2.7.7.40)	DTDP-GLUCOSE 4,6-DEHYDRATASE (EC 4.2.1.46)
NT Stop 47582	489	5880	646	3445	1202	130	866	7191	5020	4527	9627	5227	1281	6493	<del>2</del>
NT Start		5383	7	2345	2302	287	573	8351	3935	3301	8848	4448	427	7260	222
Contig.	GR00742	GR00749	GR00737	GR00718	GR00352	GR00367	GR00616	GR00367	GR00400	GR00626	VV0048	GR00002	GR00438	GR00753	GR00222
Identification Code	F RXA02596	F RXA02642	RXA02572	RXA02485	RXA01216	RXA01259	RXA02028	RXA01262	RXA01377	RXA02063	RXN00014	F RXA00014	RXA01570	RXA02666	RXA00825
Amino Acid SEQ ID NO	258	260	262	264	266	268	270	272	274	276	278	280	282	284	286
Nucleic Acid SEQ ID NO	257	259	261	263	265	267	569	271	273	275	277	279	281	283	285

## Inositol and ribitol metabolism

MYO-INOSITOL 2-DEHYDROGENASE (EC 1.1.1
3209
4219
GR00539
RXA01887
288
287

Table 1 (continued)	NT Stop Function		8838 MYO-INOSITOL-1(OR 4)-MONOPHOSPHATASE 1 (EC 3.1.3.25)	4438 MYO-INOSITOL-1(OR 4)-MONOPHOSPHATASE 1 (EC 3.1.3.25)	5504 INOSITOL MONOPHOSPHATE PHOSPHATASE	4 MYO-INOSITOL 2-DEHYDROGENASE (EC 1.1.1.18)	4 MYO-INOSITOL 2-DEHYDROGENASE (EC 1.1.1.18)	3342 MYO-INOSITOL 2-DEHYDROGENASE (EC 1.1.1.18)	4462 MYO-INOSITOL 2-DEHYDROGENASE (EC 1.1.1.18)	1977 MYO-INOSITOL 2-DEHYDROGENASE (EC 1.1.1.18)	47037 MYO-INOSITOL 2-DEHYDROGENASE (EC 1.1.1.18)	22318 MYO-INOSITOL-1-PHOSPHATE SYNTHASE (EC 5.5.1.4)	7688 MYO-INOSITOL 2-DEHYDROGENASE (EC 1.1.18)	10948 GLUCOSE-FRUCTOSE OXIDOREDUCTASE PRECURSOR (EC 1.1.99.28)	
lable	NT Start		7966	3566	6328	629	552	2338	3380	2999	48113	23406	7017	10277	931
	Contig.		VV0048	GR00002	GR00306		GR00388	GR00454	GR00454	VV0278	VV0050	VV0079	VV0028	GR10040	GR00038
	Identification Code		RXN00013	F RXA00013	RXA01099	RXN01332	F RXA01332	RXA01632	RXA01633	RXN01406	RXN01630	RXN00528	RXN03057	F RXA02902	RXA00251
	Amino Acid	SEQ ID NO	290	292	294	296	298	300	302	304	306	308	310	312	314
		SEQ ID NO													

## Utilization of sugars

Function	GLUCOSE 1-DEHYDROGENASE (EC 1.1.1.47)	GLUCONOKINASE (EC 2.7.1.12)	GLUCONOKINASE (EC 2.7.1.12)	GLUCONOKINASE (EC 2.7.1.12)	D-RIBOSE-BINDING PERIPLASMIC PROTEIN PRECURSOR	FRUCTOKINASE (EC 2.7.1.4)	FRUCTOKINASE (EC 2.7.1.4)	PERIPLASMIC BETA-GLUCOSIDASE/BETA-XYLOSIDASE PRECURSOR	(EC 3.2.1.21) (EC 3.2.1.37)	PERIPLASMIC BETA-GLUCOSIDASE/BETA-XYLOSIDASE PRECURSOR	EC 3.2.1.21) (EC 3.2.1.37)	MANNITOL 2-DEHYDROGENASE (EC 1.1.1.67)	FRUCTOSE REPRESSOR	Hypothetical Oxidoreductase (EC 1.1.1)	GLUCOSE-FRUCTOSE OXIDOREDUCTASE PRECURSOR (EC	1.99.28)	GLUCOSEFRUCTOSE OXIDOREDUCTASE PRECURSOR (EC	.1.99.28)	GLUCOSEFRUCTOSE OXIDOREDUCTASE PRECURSOR (EC	.1.99.28)	SUCROSE-6-PHOSPHATE HYDROLASE (EC 3.2.1.26)	SUCROSE-6-PHOSPHATE HYDROLASE (EC 3.2.1.26)	SUCROSE-6-PHOSPHATE HYDROLASE (EC 3.2.1.26)
		35	บ	บี	<u>о</u>	뚔	뚔	PE	<u>၁</u>	Ä	ည္			H	ਹ	-	ฮ	-	ฮ	1.1	S	S	SUC
NT Stop	13090	11114	492	1499	275	5604 4	1086	56834		1584		10520	7854	8180	z,		7050		30		z,	9	349
NT Start	12206	9633	1502	1972	1216	6557	565	58477		_		12028	6880	7035	316		6616		735		1246	725	1842
Contig.	VV0090			_		W0127	GR00240	W0009		GR00214		GR00003	GR00725	-	GR00053		0000		GR00053		GR00007	GR00615	
Identification Code	RXN02654	RXN01049	F RXA01049	F RXA01050	RXA00202	RXN00872	F RXA00872	RXN00799		F RXA00799		RXA00032	RXA02528	RXN00316	F RXA00309		RXN00310		F RXA00310		RXA00041	RXA02026	RXA02061
Amino Acid SEQ ID NO	316 318																						
Nucleic Acid SEQ ID NO	315	319	321	323	325	327	329	331		333		335	337	339	341		343		345		347	349	351

Table 1 (continued)

	MANNOSE-6-PHOSPHATE ISOMERASE (EC 5.3.1.8) MANNOSE-6-PHOSPHATE ISOMERASE (EC 5.3.1.8)	ISPHATE ISOMERASE (EC 5.3.1.8) AN BRANCHING ENZYME (EC 2.4.1.18)	AN BRANCHING ENZYME (EC 2.4.1.18)	GLYCOGEN DEBRANCHING ENZYME (EC 2.4.1.25) (EC 3.2.1.33) GLYCOGEN DEBRANCHING ENZYME (EC 2.4.1.25) (EC 3.2.1.33)	GLYCOGEN OPERON PROTEIN GLGX (EC 3.2.1)	GLYCOGEN PHOSPHORYLASE (EC 2.4.1.1)	SPHORYLASE (EC 2.4.1.1)	SPHORYLASE (EC 2.4.1.1)	SPHORYLASE (EC 2.4.1.1)	VEHORY LAVE (EC. 2.4.1.1)	GLUCOAMYLASE G1 AND G2 PRECURSOR (EC 3.2.1.3)	GLUCOSE-RESISTANCE AMYLASE REGULATOR	SE (EC 2.7.1.17)	SE (EC 2.7.1.17)	2.7.1.15)	2.7.1.15)	REPRESSOR	5-PHOSPHO-BETA-GLUCOSIDASE (EC 3.2.1.86)	DEOXYRIBOSE-PHOSPHATE ALDOLASE (EC 4.1.2.4)	1-deoxy-D-xylulose 5-phosphate reductoisomerase (EC 1.1.1)	1-deoxy-D-xylulose 5-phosphate reductoisomerase (EC 1.1.1)	-DEOXYXYLULOSE-5-PHOSPHATE SYNTHASE	-DEOXYXXLII OSE-5-PHOSPHATE SYNTHASE	1-DECKTATECEOSE-3-FRONFRAIE STNIFFASE 4-ALDEA-CLIPANOTDANSEDASE (F) 2-A 1-25	VOTRANSFERASE (EC 2.4.1.25), amvlomaltase	N-ACETYLGLUCOSAMINE-6-PHOSPHATE DEACETYLASE (EC 3.5.1.25)	SAMINE-6-PHOSPHATE DEACETYLASE (EC 3.5.1.25)	N-ACETYLGLUCOSAMINYLTRANSFERASE (EC 2.4.1)	)SAMINYLTRANSFERASE (EC 2.4.1)	SAMINYLTRANSFERASE (EC 2.4.1)	GLUCOSAMINE-6-PHOSPHATE ISOMERASE (EC 5.3.1.10)	FRUCTOSE-6-PHOSPHATE AMINOTRANSFERASE	(ISOMERIZING) (EC 2.6.1.16) JRONATE ISOMERASE (EC 5.3.1.12)	RASE. Glucuronate isomerase (EC 5.3.1.12)	URONATE ISOMERASE (EC 5.3.1.12)	JRONATE ISOMERASE, Glucuronate isomerase (EC 5.3.1.12)	GALACTOSIDE O-ACETYLTRANSFERASE (EC 2.3.1.18)	D-RIBITOL-5-PHOSPHATE CYTIDYLYLTRANSFERASE (EC 2.7.7.40)	D-RIBOSE-BINDING PERIPLASMIC PROTEIN PRECURSOR	D-RIBOSE-BINDING PERIPLASMIC PROTEIN PRECURSOR
Function	MANNOSE-6-PHO	MANNOSE-6-PHO 1,4-ALPHA-GLUCA	1,4-ALPHA-GLUCA	GLYCOGEN DEBR	GLYCOGEN OPER	GLYCOGEN PHOS	GLYCOGEN PHOS	GLYCOGEN PHOS	GLYCOGEN PHOS	ALDHA AMYI ASE (EC 3 2 4 4)	GLUCOAMYLASE	GLUCOSE-RESIST	XYLULOSE KINASE (EC 2.7.1.17)	XYLULOSE KINASE (EC 2.7.1.17)	RIBOKINASE (EC 2.7.1.15)	<b>RIBOKINASE (EC 2.7.1.15)</b>	RIBOSE OPERON REPRESSOR	6-PHOSPHO-BET/	DEOXYRIBOSE-PI	1-deoxy-D-xylulose	1-deoxy-D-xylulose	1-DEOXYXYLULO	1-DEOXYXYLULO	1-DECATATECEC	4-ALPHA-GLUCAN	N-ACETYLGLUCO	N-ACETYLGLUCO	N-ACETYLGLUCO	N-ACETYLGLUCO	N-ACETYLGLUCO	GLUCOSAMINE-6-	GLUCOSAMINE-F	(ISOMERIZING) (EC 2.6.1.16) URONATE ISOMERASE (EC (	URONATE ISOME	URONATE ISOME	URONATE ISOME	GALACTOSIDE O-	D-RIBITOL-5-PHO	D-RIBOSE-BINDIN	D-RIBOSE-BINDIN
NT Stop	1776 503	1302 1752	3985	1890	17427	16260	1346	2326	920	1207	12352	4923	49244	1118	4	2641	731	2552	5005	1103	1708	3137	1039	15/3	3828	2081	2081	33805	510	547	1279	15397	667	4	163	163	2285	6493	275	4258
NT Start	3333	595 1	1793	et	16981	14749	က	7	ო ი	15516	10517	4366	50623	ဗ	747	1739	1768	2193	9299	543	1094	1230	2 2	9/1	5927	3244	3244	35265	1157	1473	2037	17271	7	675	672	672	1611	7260	1216	2097
Contig.	VV0124 GR00398	GR00399 GR00743	GR00743	VV0184	GR00306	VV0143	GR00431	VV0318	GR00631	GK00633	GR00422	GR00539	VV0127	GR00555	GR00762	GR00778	GR00762	GR00729	GR00385	GR00030	GR00030	VV0191	GK00436	GR00480	GR00242	W0119	GR00007	W0127	GR00520	GR00529	GR00007	GR00422	VV0336	GR10013	VV0337	GR10014	GR00662	GR00753	GR00032	GR00709
Identification Code	RXN01369 F RXA01369	F RXA01373 RXA02611	RXA02612	EXN01884 F RXA01884	RXA01111	RXN01550	F RXA01550	RXN02100	F RXA02100	F KXAUZ113	RXA01478	RXA01888	RXN01927	F RXA01927	RXA02729	RXA02797	RXA02730	RXA02551	RXA01325	RXA00195	RXA00196	RXN01562	F KXA01562	P. KXAU1703	F RXA00879	RXN00043	F RXA00043	RXN01752	F RXA01839	RXA01859	RXA00042	RXA01482	RXN03179	F RXA02872	RXN03180	F RXA02873	RXA02292	RXA02666	RXA00202	RXA02440
Amino Acid SEQ ID NO	354 356	358 360	362	364 366	368	370	372	374	376	380	382	384	386	388	390	392	394	396	398	400	402	\$ 6	30 <b>5</b>	408	412	414	416	418	420	422	424	426	428	430	432	434	436	438	440	442
Nucleic Acid SEQ ID NO	353 355	357 359	361	363	367	369	371	373	375	370	381	383	385	387	389	391	393	395	397	399	401	403	403 403	404	411	413	415	417	419	421	423	425	427	429	431	433	435	437	439	144

tinued)	Function		4TDP-4-DEHYDRORHAMNOSE REDUCTASE (EC 1.1.1.133)	DIDP-4-DEHYDRORHAMNOSE REDUCIASE (EC.1.1.1.133)	DIDF-4-DEHTUKOKHAMNOVE KEDOCIAVE (EC. 1.1.1.155)	DIDT-GLUCCOS 4,9-DERTURALASE (EC. 4.2. 1.40) DIDP-GLUCOSE 4 G-DEHYDRATASE (EC. 4.2. 1.46)	ATDP-RHAMNOSYI TRANSFERASE REBE (FC 2 )	DTDP-RHAMNOSYL TRANSFERASE RFBF (EC 2)	PROTEIN ARAJ	PROTEIN ARAJ	PROTEIN ARAJ	GLUCAN ENDO-1,3-BETA-GLUCOSIDASE A1 PRECURSOR (EC 3.2.1.39)	UDP-GLUCOSE 6-DEHYDROGENASE (EC 1.1.1.22)	PULATIVE HEXULOSE-6-PHOSPHATE ISOMEKASE (EC.5)	(EC 3.2.1.21) (EC 3.2.1.37)	5-DEHYDRO-4-DEOXYGLUCARATE DEHYDRATASE (EC 4.2.1.41)	ALDOSE REDUCTASE (EC 1.1.1.21)	arabinosyl transferase subunit B (EC 2.4.2)	PHOSPHO-2-DEHYDRO-3-DEOXYHEPTONATE ALDOLASE (EC 4.1.2.15)	PUTATIVE GLYCOSYL TRANSFERASE WBIF	PUTATIVE HEXULOSE-6-PHOSPHATE ISOMERASE (EC 5)	NAGD PROTEIN	GALACTOKINASE (EC 2.7.1.6)	PHOSPHO-Z-DEHTORO-3-DEOXTHEPTONATE ALDOLASE (EC 4.1.2.13) BETA DEVOCAMINIDACE A BECTIESCO (EC 3.3.4.53)	GLUCOSE-FRUCTOSE OXIDOREDUCTASE PRECURSOR (EC	1.1.99.28)	GLUCOSE-FRUCTOSE OXIDOREDUCTASE PRECURSOR (EC	I. I.39.20) CYCLOMAI TODEXTRINGSE (FC 3.0.1.54)	CYCLOMALTODEXTRINASE (EC 3.2.1.54)	protein involved in sugar metabolism	Membrane Lipoprotein involved in sugar metabolism	Exported Protein involved in ribose metabolism	protein involved in sugar metabolism	Membrane Spanning Protein involved in metabolism of diols	Amino Acid ABC Transporter ATP-Binding Protein involved in sugar	metabolism ARC Transporter ATP-Binding Protein involved in sugar metabolism	Membrane Spanning Protein involved in sugar metabolism	Cytosolic Protein involved in sugar metabolism	Cytosolic Kinase involved in metabolism of sugars and thiamin	ABC Transporter ATP-Binding Protein involved in sugar metabolism	Membrane Spanning Protein involved in sugar metabolism	Cytosolic Protein involved in sugar metabolism Cytosolic Protein involved in sugar metabolism
Table 1 (continued)	NT Stop		42444	427	8042	21.2	6219	2022	9880	10656	11167	. 26545	∞ ;	6835	2	11489	22442	5116	38303	4750	46143	12408	21418	0640			1008		260													
Table	NT Start		41086	77	77.7	577 6103	7007	1591	10263	11147	12390	28686	289	6258		12427	23242	1679	39688	5610	47021	13274	20369	9100			<del>-</del>		1417													
	Contig.		W0009	GK00438	GR00624	GR00624	W0112	GR00098	GR00057	GR00057	GR00057	VV0135	W0063	82000	6000	VV0025	VV0102	W0181	VV0017	W0091	050000	VV0229	VV0197	VVU323			GR00549		GR00006													
	Identification Code		RXN01569	F KXA01569	F KXAUZU35	RXA00623	RXN00427	F RXA00427	RXA00327	RXA00328	RXA00329	RXN01554	RXN03015	KXN03056	000000	RXN00401	RXN02125	RXN00200	RXN01175	RXN01376	RXN01631	RXN01593	RXN00337	KX500584	RXS03215		F RXA01915	RXS03224	F RXA00038	RXC00233	RXC00236	RXC00271	RXC00338	RXC00362	RXC00412	RXC00526	RXC01004	RXC01017	RXC01021	RXC01212	KXC01306	RXC01372
	Amino Acid	SEQ ID NO	444	0 4 4 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0	2440	450	454	456	458	460	462	464	466	804	o F	472	474	476	478	480	482	4 5 4 5	486	888	492		494	496	498	200	502	<u>\$</u>	206	508	510	512	514	516	518	520	522	526 526
	Nucleic Acid	SEQ ID NO	443	445	/44/	451	453	455	457	459	461	463	465	46/	n P	471	473	475	477	479	481	483	485	784	491		493	495	497	499	501	503	505	507	60 <b>c</b>	511	513	515	517	519	521	525 525

(pa	Function	protein involved in sugar metabolism protein involved in sugar metabolism protein involved in sugar metabolism	Cytosolic Protein involved in sugar metabolism	Membrane Associated Protein involved in sugar metabolism Odrogija Drafeja javojudal ja sugar metabolism	Oytosonic motern involved in sugar metabolism protein involved in sugar metabolism	Uncharacterized protein involved in glycerol metabolism (homolog of	Drosophila rhomboid) protein involved in sugar metabolism		Function		DITRATE SYNTHASE (EC 4.1.3.7)	CITRATE LYASE BETA CHAIN (EC 4.1.3.6)	SOCITRATE DEHYDROGENASE (NADP) (EC 1.1.1.42)	SOCITRATE DEHYDROGENASE [NADP] (EC 1.1.1.42)	ACONITATE HYDRATASE (EC 4.2.1.3)	ONITATE HYDRATASE (EC 4.2.1.3)	ONITATE HYDRATASE (EC 4.2.1.3)	ONITATE HYDRATASE (EC 4.2.1.3)	2-UXUGLUTAKATE DEHYDKUGENASE ET CUMPONENT (EC 1.2.4.2)	DINTERACTION OF THE PROPERTY OF THE PROPERTY (FC.) 3.1 (FC.) OF THE PROPERTY OF THE PROPERTY (FC.) 3.1 (FC.)	SUCCINYL-COA SYNTHETASE ALPHA CHAIN (EC 6.2.1.5)	SUCCINYL-COA SYNTHETASE BETA CHAIN (EC 6.2.1.5)	L-MALATE DEHYDROGENASE (ACCEPTOR) (EC 1.1.99.16)	MALATE DEHYDROGENASE (ACCEPTOR) (EC 1.1.99.16)	MALATE DEHYDROGENASE (ACCEPTOR) (EC 1.1.99.16)	LIC ENZYME (EC 1.1.1.39)	ILIC ENZYME (EC 1.1.1.39)	MALIC ENZYME (EC 1.1.1.39)	LIC ENZYME (EC 1.1.1.39)	DIHYDROLIPOAMIDE SUCCINYLTRANSFERASE COMPONENT (E2) OF	2-OXOGLUTARATE DEHYDROGENASE COMPLEX (EC 2.3.1.61) DIHYDBOLIBOAMIDE SLICCINXI TBANSEERASE COMBONENT OF 2-	OXOGLUTARATE DEHYDROGENASE COMPLEX (EC 2.3.1.61)	oxoglutarate semialdehyde dehydrogenase (EC 1.2.1)
Table 1 (continued)	NT Stop Fu	222	.♂:	žć	ริ ธั		בֿ בֿ		NT Stop Fu		_	Ĭ	_	_								_	-				9		_	_	2-4		
le 1 (c						268					9418	1829	3372	1060	1671	1661	2151	2046	7870	)) *	3103	4009	128	9546	4179	5655	13	290	5655	583	146	<u> </u>	9922
Tab	NT Start					825			NT Start		10710	2647	5585	~	-	m	1378	1330	, ,	7	3984	5280	11307	8608	4388	4693	12539	က	4693	7	15056	3	11481
	Contig.					GR00709			Contig.		GR00641	GR00746	W0144	GR00133	<b>W0304</b>	GR00648	W0305	GR00649	GR00625	0400493	GR00206	GR00206	VV0139	GR00449	GR00474	GR00046	W0079	GR00296	GR00046	99000	30,000	2004	VV0025
	Identification Code	RXC01659 RXC01663 RXC01693	RXC01703	PXC02254	RXC02435	F RXA02435	RXC03216		Identification Code		RXA02175	RXA02621	RXN00519	F RXA00521	RXN02209	F RXA02209	RXN02213	F RXA02213	KXA02056	C*/1049	RXA00782	RXA00783	RXN01695	F RXA01615	F RXA01695	RXA00290	RXN01048	F RXA01048	F RXA00290	RXN03101	DYNOODAG	0107010	RXN00389
	Amino Acid SEQ ID NO	528 530 532	534	536	540	542	544	<u>ə</u>	Amino Acid	SEQ ID NO	546	548	550	552	554	226	558	260	562 564	204	999	568	570	572	574	976	578	580	582	584	A A	3	588
	Nucleic Acid SEQ ID NO	527 529 531	533	535 £27	539	541	543	TCA-cycle	Nucleic Acid	SEQ ID NO	545	547	549	551	553	555	557	559	561 563	Sec	565	267	569	571	573	575	277	679	581	583	787	2	587

### Table 1 (continued)

Glyoxylate bypass

Function	ISOCITRATE LYASE (EC 4.1.3.1) ISOCITRATE LYASE (EC 4.1.3.1) MALATE SYNTHASE (EC 4.1.3.2) MALATE SYNTHASE (EC 4.1.3.2) GLYOXYLATE-INDUCED PROTEIN GLYOXYLATE-INDUCED PROTEIN	
NT Stop	18365 1773 22475 1663 3958 2430	
NT Start	19708 478 20259 3798 3209	
Contig.	VV0176 GR00699 VV0176 GR00700 GR00304 GR00539	
Identification Code	RXN02399 F RXA02399 RXN02404 F RXA02404 RXA01089 RXA01886	
Amino Acid SEQ ID NO	590 592 594 598 600	
Nucleic Acid SEQ ID NO	589 591 595 597 599	

## Methylcitrate-pathway

Function	2-methylisocitrate synthase (EC 5.3.3)	lisocitrate syr	2-methylcitrate synthase (EC 4.1.3.31)	2-methylisocitrate synthase (EC 5.3.3)	2-methylcitrate synthase (EC 4.1.3.31)	methylisocitrate lyase (EC 4.1.3.30)	methylisocitrate lyase (EC 4.1.3.30)	LACTOYLGLUTATHIONE LYASE (EC 4.4.1.5)			
NT Stop	1576 4	1576	2773	6017	90	S	ည	764	1815	1902	9979
NT Start	3087 978	1983	3069	4647	7	415	607	1906	901	2120	9230
Contig.	VV0092	GR00130	GR00131	GR00300	W0141	GR00668	GR00669	GR00671	W0141	GR00671	GR00003
Identification Code	RXN03117 F RXA00406	F RXA00514	RXA00518	RXA01077	EXN03144	F RXA02322	RXA02329	RXA02332	RXN02333	F RXA02333	RXA00030
Amino Acid SEQ ID NO	602	909	610	612	614	616	618	620	622	624	626
Nucleic Acid SEQ ID NO	600	603 603	607	609	611	613	615	617	619	621	623

## Methyl-Malonyl-CoA-Mutases

Function		METHYLMALONYL-COA MUTASE ALPHA-SUBUNIT (EC 5.4.99.2)	METHYLMALONYL-COA MUTASE ALPHA-SUBUNIT (EC 5.4.99.2)	METHYLMALONYL-COA MUTASE BETA-SUBUNIT (EC 5.4.99.2)
NT Stop			2	
NT Start		9849	2002	3856
Contig.		VV0167	GR00023	GR00023
Identification Code		RXN00148	F RXA00148	RXA00149
Amino Acid	SEQ ID NO	628	630	632
Nucleic Acid	SEQ ID NO	625	627	629

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Tabl	

Function	PHOSPHOGLYCOLATE PHOSPHATASE (EC 3.1.3.18) PHOSPHOGLYCOLATE PHOSPHATASE (EC 3.1.3.18) PHOSPHOGLYCOLATE PHOSPHATASE (EC 3.1.3.18) PHOSPHOGLYCOLATE PHOSPHATASE (EC 3.1.3.18)		Function	CYTOCHROME D UBIQUINOL OXIDASE SUBUNIT I (EC 1.10.3) CYTOCHROME D UBIQUINOL OXIDASE SUBUNIT I (EC 1.10.3)	CYTOCHROME D UBIQUINOL OXIDASE SUBUNIT (EC 1.10.3) CYTOCHROME C-TYPE BIOGENESIS PROTEIN CODA	CYTOCHROME C-TYPE BIOGENESIS PROTEIN CCDA	CYTOCHROME D UBIQUINOL OXIDASE SUBUNIT II (EC 1.10.3) CYTOCHROME C OXIDASE DOI YPEPTIDE I /EC 1.9.3.1)	CYTOCHROME C OXIDASE SUBUNIT I (EC 1.9.3.1)	CYTOCHROME C OXIDASE POLYPEPTIDE I (EC 1.9.3.1)	CYTOCHROME C OXIDASE POLYPEPTIDE I (EC 1.9.3.1)	CYTOCHROME C OXIDASE POLYPEPTIDE II (EC 1.9.3.1)	CYTOCHROME C OXIDASE POLYPEPTIDE I (EC 1.9.3.1) RIESKE IRON, SLII ELIB DROTEIN	PROBABLE CYTOCHROME C OXIDASE ASSEMBLY FACTOR	CYTOCHROME AA3 CONTROLLING PROTEIN	FERREDOXIN	FERREDOXIN	FERREDOXIN VI	ELECTRON TRANSFER FLAVOPROTEIN ALPHA-SUBUNIT	ELECTRON TRANSFER FLAVOPROTEIN BETA-SUBUNIT	NADH DEHYDROGENASE I CHAIN L (EC 1.6.5.3)	NADH DEHYDROGENASE I CHAIN L (EC 1.6.5.3)	NADH DEHYDROGENASE I CHAIN M (EC 1.6.5.3)	NADH DEHYDROGENASE I CHAIN M (EC 1.6.5.3)	NADH DEHYDROGENASE I CHAIN L (EC 1.6.5.3)	NADH DEHYDROGENASE I CHAIN L (EC 1.6.5.3)	NADH-UBIQUINONE OXIDOREDUCTASE CHAIN 2	NADH-UBIQUINONE OXIDOREDUCTASE 39 KD SUBUNIT PRECURSOR	(兵し 1.6.5.3) (モレ 1.5.99.3)
NT Stop	27532 6 3264 14643		NT Stop	812 11890	218 6	435	6 29567	4	601	1334	8415	10063 12248	8542	12497	1519	122	2315	24015	24998	9056	1869	7113	3017	2120	3406	£	46287	
NT Start	26879 344 3956 14236		NT Start	2350 11753	212	773	31222	288	1449	1945	7339	9413	7613	13534	1199	436	2632	24965	25783	11299	121	8642	2253	က	2552	846	44824	
Contig.	VV0197 GR00055 GR00645 VV0124		Contig.	VV0174 GR00008	GR00494 GR00082	GR00083	GR00494 VV0084	GR00550	GR00717	GR00717	GR00639	GR00639	GR00763	GR00763	GR00355	GR00532	GR00179	GR00032	GR00032	W0192	GR00160	W0192	GR00160	GR00249	GR00247	GR00182	00000	
Identification Code	RXN00317 F RXA00317 RXA02196 RXN02461		Identification Code	RXN01744 F RXA00055	F KXA01744	RXA00385	RXA01743 RXN02480	F RXA01919	F RXA02480	F RXA02481	RXA02140	RXA02142 RXA02144	RXA02740	RXA02743	RXA01227	EXA01865	EXA00680	RXA00224	RXA00225	RXN00606	F RXA00606	RXN00595	F RXA00608	RXA00913	RXA00909	RXA00700	RXN00483	
Amino Acid	634 636 638 640	hain	Amino Acid SEQ ID NO	642 644	648 648	650	652 654	656	658	099	662	664 666	668	029	672	674	678 678	089	682	684	989	688	069	692	694	969	869	
Nucleic Acid	631 635 637 639	Redox Chain	Nucleic Acid SEQ ID NO	643	647 647	649	651 653	655	657	659	661	663 665	299	699	671	673	6/5 677	679	681	683	685	289	689	169	693	695	269	

		NADH-UBIQUINONE OXIDOREDUCTASE 39 KD SUBUNIT PRECURSOR FEC 1 6.5.3 (FC 1 6.99.3)	DUCTASE	(6.5.5)	.6.5.5)	(EC 1.6.99)	(EC 1.6.99)	ON-SULFUR PROTEIN (EC 1.3.99.1)	A (EC 1.6.5.3)		9.3)	1	1A CHAIN (EC 1.2.1.2)			TEIN CCSA	O	ilar to cytochrome c biogenesis			TERIDINE REDUCTASE (EC		C 2.5.1.18)	GLUTATHIONE-DEPENDENT FORMALDEHYDE DEHYDROGENASE (EC	•	ome c oxidoreductase	A (EC 1.6.5.3)	<b>TASE CHAIN 4 (EC 1.6.5.3)</b>			<u>~</u>
tinued)	Function	NADH-UBIQUINONE OXIDOREDUC	NADH-DEPENDENT FMN OXYDOREDUCTASE	QUINONE OXIDOREDUCTASE (EC	QUINONE OXIDOREDUCTASE (EC 1.6.5.5)	NADPH-FLAVIN OXIDOREDUCTASE (EC 1.6.99)	NADPH-FLAVIN OXIDOREDUCTASE	SUCCINATE DEHYDROGENASE IRON-SULFUR PROTEIN (EC 1.3.99.1)	NADH DEHYDROGENASE I CHAIN M (EC 1.6.5.3)	Hydrogenase subunits	NADH DEHYDROGENASE (EC 1.6.99.3)	DEHYDROGENASE	FORMATE DEHYDROGENASE ALPHA CHAIN (EC 1.2.1.2)	FDHD PROTEIN	FDHD PROTEIN	CYTOCHROME C BIOGENESIS PROTEIN CCSA	essential protein similar to cytochrome c	RESC PROTEIN, essential protein similar to cytochrome c biogenesis	protein	putative cytochrome oxidase	FLAVOHEMOPROTEIN / DIHYDROPTERIDINE REDUCTASE (EC	1.6.99.7) FI AVOHEMOPROTEIN	GLUTATHIONE S-TRANSFERASE (EC 2,5,1,18)	GLUTATHIONE-DEPENDENT FORM	1.2.1.1)	QCRC PROTEIN, menaquinol:cytochrome c oxidoreductase	NADH DEHYDROGENASE I CHAIN M (EC 1.6.5.3)	NADH-UBIQUINONE OXIDOREDUCTASE CHAIN 4 (EC 1.6.5.3)	Hypothetical Oxidorductase	Hypothetical Oxidoreductase	Hypothetical Oxidoreductase (EC 1.1.1)
Table 1 (continued)	NT Stop	20569	547	1636	8620	10788	7160	865	368	1259	2	817	271	5197	407	3091	299	S.		2847	6229	3176	3373	3134		11025	4	33063	2794	849	4010
Table	NT Start	19106	1035	2646	9585	9922	6339	1611	1273	ဗ	955	7	2556	6111	1291	2081	696	514		1876	2095	2019	2297	2031		10138	405	32683	3552	1784	4633
	Contig.	GR00119	GR00427	GR00046	GR00763	W0101	GR00731	GR00380	VV0058	GR00248	W0117	GR00543	GR00183	VV0005	GR00184	VV0025	GR00085	GR00084		GR00259	W0101	GR00731	GR00408	GR00214		GR00639	VV0058	VV0176	VV0317	VV0302	VV0101
	Identification Code	F RXA00483	RXA01534	RXA00288	RXA02741	RXN02560	F RXA02560	RXA01311	RXN03014	F RXA00910	RXN01895	F RXA01895	RXA00703	RXN00705	F RXA00705	RXN00388	F RXA00388	F RXA00386		RXA00945	RXN02556	F RXA02556	RXA01392	RXA00800		RXA02143	RXN03096	RXN02036	RXN02765	RXN02206	RXN02554
	Amino Acid SEQ ID NO	200	702	704	206	708	710	712	714	716	718	720	722	724	726	728	730	732		734	736	738	740	742		744	746	748	750	752	754
	Nucleic Acid SEQ ID NO	669	701	703	705	707	209	711	713	715	717	719	721	723	725	727	729	731		733	735	737	739	741		743	745	747	749	751	753

### ATP-Synthase

Function		ATP SYNTHASE A CHAIN (EC 3.6.1.34)	ATP SYNTHASE A CHAIN (EC 3.6.1.34)	ATP SYNTHASE ALPHA CHAIN (EC 3.6.1.34	ATP SYNTHASE BETA CHAIN (EC 3.6.1.34)	ATP SYNTHASE BETA CHAIN (EC 3.6.1.34)	ATP SYNTHASE BETA CHAIN (EC 3.6.1.34)
NT Stop		461	1155	2315	3832	755	3993
NT Start		1270	394	675	5280	15	3355
Contig.		VV0121	GR00345	GR00344	W0175	GR00343	GR00344
Identification Code		RXN01204	F RXA01204	RXA01201	RXN01193	F RXA01193	F RXA01203
Amino Acid	SEQ ID NO	756	758	760	762	764	99/
	SEO ID NO						

unea)	Function		ATP SYNTHASE C CHAIN (EC 3.6.1.34)	ATP SYNTHASE C CHAIN (EC 3.6.1.34)	ATP SYNTHASE DELTA CHAIN (EC 3.6.1.34)	ATP SYNTHASE EPSILON CHAIN (EC 3.6.1.34)	ATP SYNTHASE GAMMA CHAIN (EC 3.6.1.34)	ATP-BINDING PROTEIN		Function	CYTOCHROME P450 116 (EC 1.14) Hypothetical Cytochrome c Biogenesis Protein
lable 1 (continued	NT Stop		85	318	610	1141	3349	3274		NT Stop	28581 2004
	NT Start		324	139	2	770	2375	4923		NT Start	29864 1150
	Contig.		VV0121	GR00802	GR00344	GR00343	GR00344	0600//		Contig.	VV0005 VV0025
	Identification Code		RXN02821	F RXA02821	RXA01200	RXA01194	RXA01202	RXN02434	oolism	Identification Code	RXN00684 RXN00387
	Amino Acid	SEQ ID NO	768	077	772	774	216	778	Cytochrome metabolism	Amino Acid SEQ ID NO	780 782
	Nucleic Acid	SEQ ID NO	792	69/	171	773	775	717	Cytochro	Nucleic Acid SEQ ID NO	779 781

		TABLE 2 – Excluded Genes	ded Genes
GenBank <sup>TM</sup> Accession No.	Gene Name	Gene Function	Reference
A09073	gdd	Phosphoenol pyruvate carboxylase	Bachmann, B. et al. "DNA fragment coding for phosphoenolpyruvat corboxylase, recombinant DNA carrying said fragment, strains carrying the recombinant DNA and method for producing L-aminino acids using said strains," Patent: EP 0358940-A 3 03/21/90
A45579, A45581, A45583, A45585		Threonine dehydratase	Moeckel, B. et al. "Production of L-isoleucine by means of recombinant micro-organisms with deregulated threonine dehydratase," Patent: WO 9519442-A 5 07/20/95
AB003132	murC; ftsQ; ftsZ		Kobayashi, M. et al. "Cloning, sequencing, and characterization of the ftsZ gene from coryneform bacteria," Biochem. Biophys. Res. Commun., 236(2):383-388 (1997)
AB015023	murC; ftsQ		Wachi, M. et al. "A murC gene from Coryneform bacteria," Appl. Microbiol. Biotechnol., 51(2):223-228 (1999)
AB018530	dtsR		Kimura, E. et al. "Molecular cloning of a novel gene, dtsR, which rescues the detergent sensitivity of a mutant derived from <i>Brevibacterium lactofermentum</i> ," <i>Biosci. Biotechnol. Biochem.</i> , 60(10):1565-1570 (1996)
AB018531	dtsR1; dtsR2		
AB020624	murl	D-glutamate racemase	
AB023377	tkt	transketolase	
AB024708	gltB; gltD	Glutamine 2-oxoglutarate aminotransferase large and small subunits	
AB025424	acn	aconitase	
AB027714	rep	Replication protein	
AB027715	rep; aad	Replication protein; aminoglycoside adenyltransferase	
AF005242	argC	N-acetylglutamate-5-semialdehyde dehydrogenase	
AF005635	glnA	Glutamine synthetase	
AF030405	hisF	cyclase	
AF030520	argG	Argininosuccinate synthetase	
AF031518	argF	Ornithine carbamolytransferase	
AF036932	aroD	3-dehydroquinate dehydratase	
AF038548	pyc	Pyruvate carboxylase	

		Table 2 (continued)	nued)
AF038651	dciAE; apt; rel	protein; a	Wehmeier, L. et al. "The role of the Corynebacterium glutamicum rel gene in (p)ppGpp metabolism," <i>Microbiology</i> , 144:1853-1862 (1998)
AF041436	argR	Arginine repressor	
AF045998	impA	Inositol monophosphate phosphatase	
AF048764	argH	Argininosuccinate lyase	
AF049897	argC; argJ; argB;	N-acetylglutamylphosphate reductase;	
********	argD; argF; argR;	ornithine acetyltransferase; N-	
	argG; argH	acetylglutamate kinase; acetylomithine	
		transminase; ornithine	
		carbamoyltransferase; arginine repressor;	
*******		argininosuccinate synthase;	
		argininosuccinate lyase	
AF050109	inhA	Enoyl-acyl carrier protein reductase	
AF050166	hisG	ATP phosphoribosyltransferase	
AF051846	hisA	Phosphoribosylformimino-5-amino-1-	
		phosphoribosyl-4-imidazolecarboxamide	
		Isomerase	
AF052652	metA	Homoserine O-acetyltransferase	Park, S. et al. "Isolation and analysis of metA, a methionine biosynthetic gene encoding homoserine acetyltransferase in Corynebacterium glutamicum," Mol. Coll. 8(3):286-204 (1908)
AF053071	aroB	Dehydroquinate synthetase	
AF060558	HisH	Glutamine amidotransferase	
AF086704	hisE	Phosphoribosyl-ATP- pyrophosphohydrolase	
AF114233	aroA	5-enolpyruvylshikimate 3-phosphate synthase	
AF116184	panD	L-aspartate-alpha-decarboxylase precursor	Dusch, N. et al. "Expression of the Corynebacterium glutamicum panD gene encoding L-aspartate-alpha-decarboxylase leads to pantothenate overproduction in Escherichia coli," <i>Appl. Environ. Microbiol.</i> , 65(4)1530-1539 (1999)
AF124518	aroD; aroE	3-dehydroquinase; shikimate dehydrogenase	
AF124600	aroC; aroK; aroB;	Chorismate synthase; shikimate kinase; 3-	
	рерQ	dehydroquinate synthase; putative cytoplasmic peptidase	
AF145897	inhA		
AF145898	inhA		

	Table 2 (continued)	nued)
ectP	Transport of ectoine, glycine betaine, proline	Peter, H. et al. "Corynebacterium glutamicum is equipped with four secondary carriers for compatible solutes: Identification, sequencing, and characterization of the proline/ectoine uptake system, ProP, and the ectoine/proline/glycine betaine carrier, EctP," J. Bacteriol., 180(22):6005-6012 (1998)
dapD	Tetrahydrodipicolinate succinylase (incomplete)	Wehrmann, A. et al. "Different modes of diaminopimelate synthesis and their role in cell wall integrity: A study with Corynebacterium glutamicum," J. Bacteriol., 180(12):3159-3165 (1998)
ppc; secG; amt; ocd; soxA	Phosphoenolpyruvate-carboxylase; ?; high affinity ammonium uptake protein; putative ornithine-cyclodecarboxylase; sarcosine oxidase	
ftsY, glnB, glnD; srp; amtP	Involved in cell division; PII protein; uridylyltransferase (uridylyl-removing enzmye); signal recognition particle; low affinity ammonium uptake protein	Jakoby, M. et al. "Nitrogen regulation in Corynebacterium glutamicum; Isolation of genes involved in biochemical characterization of corresponding proteins," FEMS Microbiol, 173(2):303-310 (1999)
cat	Chloramphenicol aceteyl transferase	
овш	L-malate: quinone oxidoreductase	Molenaar, D. et al. "Biochemical and genetic characterization of the membrane-associated malate dehydrogenase (acceptor) from Corynebacterium glutamicum," Eur. J. Biochem., 254(2):395-403 (1998)
ugu	NADH dehydrogenase	
porA	Porin	Lichtinger, T. et al. "Biochemical and biophysical characterization of the cell wall porin of Corynebacterium glutamicum: The channel is formed by a low molecular mass polypeptide," <i>Biochemistry</i> , 37(43):15024-15032 (1998)
	Transposable element IS31831	Vertes et al. "Isolation and characterization of IS31831, a transposable element from Corynebacterium glutamicum," Mol. Microbiol., 11(4):739-746 (1994)
odhA	2-oxoglutarate dehydrogenase	Usuda, Y. et al. "Molecular cloning of the Corynebacterium glutamicum (Brevibacterium lactofermentum AJ12036) odhA gene encoding a novel type of 2-oxoglutarate dehydrogenase," <i>Microbiology</i> , 142:3347-3354 (1996)
hdh; hk	Homoserine dehydrogenase; homoserine kinase	Katsumata, R. et al. "Production of L-thereonine and L-isoleucine," Patent: JP 1987232392-A 1 10/12/87
	Upstream of the start codon of homoserine kinase gene	Katsumata, R. et al. "Production of L-thereonine and L-isoleucine," Patent: JP 1987232392-A 2 10/12/87
	Tryptophan operon	
tpL; tpE	Leader peptide; anthranilate synthase	Matsui, K. et al. "Tryptophan operon, peptide and protein coded thereby, utilization of tryptophan operon gene expression and production of tryptophan," Patent: JP 1987244382-A 1 10/24/87

Table 2 (continued)	ator regions of	uthirzation of tryptophan operon peron gene expression and production of tryptophan," Patent: JP 1987244382-A 1 10/24/87	Biotin-synthase Hatakeyama, K. et al. "DNA fragment containing gene capable of coding biotin synthetase and its utilization," Patent: JP 1992278088-A 1 10/02/92	Diamino pelargonic acid aminotransferase Kohama, K. et al. "Gene coding diaminopelargonic acid aminotransferase and desthiobiotin synthetase and its utilization," Patent: JP 1992330284-A 1 11/18/92	Desthiobiotinsynthetase Kohama, K. et al. "Gene coding diaminopelargonic acid aminotransferase and desthiobiotin synthetase and its utilization," Patent: JP 1992330284-A 1 11/18/92	Flavum aspartase Kurusu, Y. et al. "Gene DNA coding aspartase and utilization thereof," Patent: JP 1993030977-A 1 02/09/93	Isocitric acid lyase Katsumata, R. et al. "Gene manifestation controlling DNA," Patent: JP 1993056782-A 3 03/09/93	Isocitric acid lyase N-terminal fragment Katsumata, R. et al. "Gene manifestation controlling DNA," Patent: JP 1993056782-A 3 03/09/93	Prephenate dehydratase Sotouchi, N. et al. "Production of L-phenylalanine by fermentation," Patent: JP 1993076352-A 2 03/30/93	Aspartokinase Fugono, N. et al. "Gene DNA coding Aspartokinase and its use," Patent: JP 1993184366-A 1 07/27/93	Dihydro-dipichorinate synthetase Hatakeyama, K. et al. "Gene DNA coding dihydrodipicolinic acid synthetase and its use," Patent: JP 1993184371-A 1 07/27/93	Diaminopimelic acid dehydrogenase Kobayashi, M. et al. "Gene DNA coding Diaminopimelic acid dehydrogenase and its use," Patent: JP 1993284970-A 1 11/02/93	Threonine synthase Kohama, K. et al. "Gene DNA coding threonine synthase and its use," Patent:  JP 1993284972-A I 11/02/93	Prephenate dehydratase Kikuchi, T. et al. "Production of L-phenylalanine by fermentation method,"  Patent: JP 1993344881-A 1 12/27/93	Mutated Prephenate dehydratase Kikuchi, T. et al. "Production of L-phenylalanine by fermentation method,"  Patent: JP 1993344881-A 1 12/27/93	Acetohydroxy acid synthetase Inui, M. et al. "Gene capable of coding Acetohydroxy acid synthetase and its 'use," Patent: JP 1993344893-A 1 12/27/93		Mutated aspartokinase alpha subunit Sugimoto, M. et al. "Mutant aspartokinase gene," patent: JP 1994062866-A 1 03/08/94	
	E01377		E03937	E04040	E04041	E04307	E04376	E04377	E04484	E05108	E05112	E05776	E05779	E06110	E06111	E06146	E06825	E06826	

Table 2 (contin	Mutated aspartokinase alpha subunit	secY	Aspartokinase	Feedback inhibition-released Aspartokinase		Acetohydroxy-acid isomeroreductase Inui, M. et al. "Gene DNA coding acetohydroxy acid isomeroreductase,"  Patent: JP 1994277067-A 1 10/04/94	Зээs	FT aminotransferase and desthiobiotin synthetase promoter region	Biotin synthetase Hatakeyama, K. et al. "DNA fragment having promoter function in coryneform bacterium," Patent: JP 1995031476-A 1 02/03/95	Aspartase	Dihydrodipicolinate reductase	Diaminopimelic acid decarboxylase	Serine hydroxymethyltransferase	0, transposase Moriya, M. et al. "Amplification of gene using artificial transposon," Patent: 9, JP 1997070291-A 03/18/97 8			aspartokinase Moriya, M. et al. "Amplification of gene using artificial transposon," Patent:  JP 1997070291-A 03/18/97	3 Dihydrodipicolinic acid reductase Moriya, M. et al. "Amplification of gene using artificial transposon," Patent: JP 1997070291-A 03/18/97
	E06827	E07701	E08177	E08178,	E08179, E08180, E08181, E08182	E08232	E08234	E08643	E08646	E08649	E08900	E08901	E12594	E12760, E12759, E12758	E12764	E12767	E12770	E12773

llvA		Glucose-6-phosphate dehydrogenase Hatak  Threonine dehydratase Of coo	Hatakeyama, K. et al. "Glucose-6-phosphate dehydrogenase and DNA capable of coding the same," Patent: JP 1997224661-A 1 09/02/97  Moeckel, B. et al. "Functional and structural analysis of the threonine dehydratase of Cornebacterium olutamicum." J Racierial, 174,8065,8077
EC 4	EC 4.2.1.15	3-deoxy-D-arabinoheptulosonate-7- phosphate synthase	(1992) Chen, C. et al. "The cloning and nucleotide sequence of Corynebacterium glutamicum 3-deoxy-D-arabinoheptulosonate-7-phosphate synthase gene," FEMS Microbiol. Lett. 107:223-230 (1993)
llvB;	IIvB; iIvN; iIvC	Acetohydroxy acid synthase large subunit; Acetohydroxy acid synthase small subunit; Acetohydroxy acid isomeroreductase	Keilhauer, C. et al. "Isoleucine synthesis in Corynebacterium glutamicum: molecular analysis of the ilvB-ilvN-ilvC operon," J. Bacteriol., 175(17):5595-5603 (1993)
PtsM		Phosphoenolpyruvate sugar phosphotransferase	Fouet, A et al. "Bacillus subtilis sucrose-specific enzyme II of the phosphotransferase system: expression in Escherichia coli and homology to enzymes II from enteric bacteria," PNAS USA, 84(24):8773-8777 (1987); Lee, J.K. et al. "Nucleotide sequence of the gene encoding the Corynebacterium glutamicum mannose enzyme II and analyses of the deduced protein sequence," FEMS Microbiol. Lett., 119(1-2):137-145 (1994)
aceB	8	Malate synthase	Lee, H-S. et al. "Molecular characterization of aceB, a gene encoding malate synthase in Corynebacterium glutamicum," J. Microbiol. Biotechnol., 4(4):256-263 (1994)
		Pyruvate kinase	Jetten, M. S. et al. "Structural and functional analysis of pyruvate kinase from Corynebacterium glutamicum," <i>Appl. Environ. Microbiol.</i> , 60(7):2501-2507 (1994)
aceA	A	Isocitrate lyase	
dtxr		Diphtheria toxin repressor	Oguiza, J.A. et al. "Molecular cloning, DNA sequence analysis, and characterization of the Corynebacterium diphtheriae dtxR from Brevibacterium lactofermentum," J. Bacteriol., 177(2):465-467 (1995)
		Prephenate dehydratase	Follettie, M.T. et al. "Molecular cloning and nucleotide sequence of the Corynebacterium glutamicum pheA gene," J. Bacteriol., 167:695-702 (1986)
5S	SS rRNA		Park, Y-H. et al. "Phylogenetic analysis of the coryneform bacteria by 56 rRNA sequences," J. Bacteriol, 169:1801-1806 (1987)
trpE	ш	Anthranilate synthase, 5' end	Sano, K. et al. "Structure and function of the trp operon control regions of Brevibacterium lactofermentum, a glutamic-acid-producing bacterium," Gene, 52:191-200 (1987)
trpA	A	Tryptophan synthase, 3'end	Sano, K. et al. "Structure and function of the trp operon control regions of Brevibacterium lactofermentum, a glutamic-acid-producing bacterium," Gene, 52:191-200 (1987)

		Table 2 (continued)	(panu
M25819		Phosphoenolpyruvate carboxylase	O'Regan, M. et al. "Cloning and nucleotide sequence of the Phosphoenolpyruvate carboxylase-coding gene of Corynebacterium glutamicum ATCC13032," Gene, 77(2):237-251 (1989)
M85106		23S rRNA gene insertion sequence	Roller, C. et al. "Gram-positive bacteria with a high DNA G+C content are characterized by a common insertion within their 23S rRNA genes," J. Gen. Microbiol., 138:1167-1175 (1992)
M85107, M85108		23S rRNA gene insertion sequence	Roller, C. et al. "Gram-positive bacteria with a high DNA G+C content are characterized by a common insertion within their 23S rRNA genes," J. Gen. Microbiol., 138:1167-1175 (1992)
M89931	aecD; bmQ; yhbw	Beta C-S Iyase; branched-chain amino acid uptake carrier; hypothetical protein yhbw	Rossol, I. et al. "The Corynebacterium glutamicum aecD gene encodes a C-S lyase with alpha, beta-elimination activity that degrades aminoethylcysteine," <i>J. Bacteriol.</i> , 174(9):2968-2977 (1992); Tauch, A. et al. "Isoleucine uptake in Corynebacterium glutamicum ATCC 13032 is directed by the bmQ gene product," <i>Arch. Microbiol.</i> , 169(4):303-312 (1998)
859299	<b>d</b> i	Leader gene (promoter)	Herry, D.M. et al. "Cloning of the trp gene cluster from a tryptophan-hyperproducing strain of Corynebacterium glutamicum: identification of a mutation in the trp leader sequence," <i>Appl. Environ. Microbiol.</i> , 59(3):791-799 (1993)
U11545	πр	Anthranilate phosphoribosyltransferase	O'Gara, J.P. and Dunican, L.K. (1994) Complete nucleotide sequence of the Corynebacterium glutamicum ATCC 21850 tpD gene." Thesis, Microbiology Department, University College Galway, Ireland.
U13922	cgliM; cgliR; clgliR	Putative type II 5-cytosoine methyltransferase; putative type II restriction endonuclease; putative type I or type III restriction endonuclease	Schafer, A. et al. "Cloning and characterization of a DNA region encoding a stress-sensitive restriction system from Corynebacterium glutamicum ATCC 13032 and analysis of its role in intergeneric conjugation with Escherichia coli," J. Bacteriol., 176(23):7309-7319 (1994); Schafer, A. et al. "The Corynebacterium glutamicum cglIM gene encoding a 5-cytosine in an McrBC-deficient Escherichia coli strain," Gene, 203(2):95-101 (1997)
U14965 U31224	recA		Ankri, S. et al. "Mutations in the Corynebacterium glutamicumproline
			biosynthetic pathway: A natural bypass of the proA step," J. Bacteriol., 178(15):4412-4419 (1996)
U31225	proC	L-proline: NADP+ 5-oxidoreductase	Ankri, S. et al. "Mutations in the Corynebacterium glutamicumproline biosynthetic pathway: A natural bypass of the proA step," J. Bacteriol., 178(15):4412-4419 (1996)
U31230	obg; proB; unkdh	?;gamma glutamyl kinase;similar to D- isomer specific 2-hydroxyacid dehydrogenases	Ankri, S. et al. "Mutations in the Corynebacterium glutamicumproline biosynthetic pathway: A natural bypass of the proA step," J. Bacteriol., 178(15):4412-4419 (1996)

		Table 2 (continued)	ned)
U31281	bioB	Biotin synthase	Serebriiskii, I.G., "Two new members of the bio B superfamily: Cloning, sequencing and expression of bio B genes of Methylobacillus flagellatum and Corynebacterium glutamicum," Gene, 175:15-22 (1996)
U35023	thtR; accBC	Thiosulfate sulfurtransferase; acyl CoA carboxylase	Jager, W. et al. "A Corynebacterium glutamicum gene encoding a two-domain protein similar to biotin carboxylases and biotin-carboxyl-carrier proteins," <i>Arch. Microbiol.</i> , 166(2);76-82 (1996)
U43535	cmr	Multidrug resistance protein	Jager, W. et al. "A Corynebacterium glutamicum gene conferring multidrug resistance in the heterologous host Escherichia coli," J. Bacteriol., 179(7):2449-2451 (1997)
U43536	clpB	Heat shock ATP-binding protein	
U53587	aphA-3	3'5"-aminoglycoside phosphotransferase	
U89648		Corynebacterium glutamicum unidentified sequence involved in histidine biosynthesis, partial sequence	
X04960	trpA; trpB; trpC; trpD; trpE; trpG; trpL	Tryptophan operon	Matsui, K. et al. "Complete nucleotide and deduced amino acid sequences of the Brevibacterium lactofermentum tryptophan operon," Nucleic Acids Res., 14(24):10113-10114 (1986)
X07563	lys A	DAP decarboxylase (meso-diaminopimelate decarboxylase, EC 4.1.1.20)	Yeh, P. et al. "Nucleic sequence of the JysA gene of Corynebacterium glutamicum and possible mechanisms for modulation of its expression," Mol. Gen. Genet., 212(1):112-119 (1988)
X14234	EC 4.1.1.31	Phosphoenolpyruvate carboxylase	Eikmanns, B.J. et al. "The Phosphoenolpyruvate carboxylase gene of Corynebacterium glutamicum: Molecular cloning, nucleotide sequence, and expression," <i>Mol. Gen. Genet.</i> , 218(2):330-339 (1989); Lepiniec, L. et al. "Sorghum Phosphoenolpyruvate carboxylase gene family: structure, function and molecular evolution," <i>Plant. Mol. Biol.</i> , 21 (3):487-502 (1993)
X17313	fda	Fructose-bisphosphate aldolase	Von der Osten, C.H. et al. "Molecular cloning, nucleotide sequence and fine- structural analysis of the Corynebacterium glutamicum fda gene: structural comparison of C. glutamicum fructose-1, 6-biphosphate aldolase to class I and class II aldolases," Mol. Microbiol.
X53993	dapA	L-2, 3-dihydrodipicolinate synthetase (EC 4.2.1.52)	Bonnassie, S. et al. "Nucleic sequence of the dapA gene from Corynebacterium glutamicum," Nucleic Acids Res., 18(21):6421 (1990)
X54223		AttB-related site	Cianciotto, N. et al. "DNA sequence homology between att B-related sites of Corynebacterium diphtheriae, Corynebacterium ulcerans, Corynebacterium glutamicum, and the attP site of lambdacorynephage," FEMS. Microbiol, Lett., 66:299-302 (1990)
X54740	argS; lysA	Arginyl-tRNA synthetase; Diaminopimelate decarboxylase	Marcel, T. et al. "Nucleotide sequence and organization of the upstream region of the Corynebacterium glutamicum lysA gene," Mol. Microbiol., 4(11):1819-1830 (1990)

		Table 2 (continued)	nied)
X55994	troL: troE	Putative leader peptide: anthranilate	Heery D.M. et al. "Nucleotide sequence of the Corynebacterium Plutamicum
		synthase component 1	trpE gene," Nucleic Acids Res., 18(23):7138 (1990)
X56037	thrC	Threonine synthase	Han, K.S. et al. "The molecular structure of the Corynebacterium glutamicum threonine synthase gene," Mol. Microbiol., 4(10):1693-1702 (1990)
X56075	attB-related site	Attachment site	Cianciotto, N. et al. "DNA sequence homology between att B-related sites of Corynebacterium diphtheriae, Corynebacterium ulcerans, Corynebacterium glutamicum, and the attP site of lambdacorynephage," FEMS. Microbiol, Lett., 66:299-302 (1990)
X57226	lysC-alpha; lysC-beta; asd	Aspartokinase-alpha subunit; Aspartokinase-beta subunit; aspartate beta semialdehyde dehydrogenase	Kalinowski, J. et al. "Genetic and biochemical analysis of the Aspartokinase from Corynebacterium glutamicum," Mol. Microbiol., 5(5):1197-1204 (1991); Kalinowski, J. et al. "Aspartokinase genes lysC alpha and lysC beta overlap and are adjacent to the aspertate beta-semialdehyde dehydrogenase gene asd in Corynebacterium glutamicum," Mol. Gen. Genet., 224(3):317-324 (1990)
X59403	gap;pgk; tpi	Glyceraldehyde-3-phosphate; phosphoglycerate kinase; triosephosphate isomerase	Eikmanns, B.J. "Identification, sequence analysis, and expression of a Corynebacterium glutamicum gene cluster encoding the three glycolytic enzymes glyceraldehyde-3-phosphate dehydrogenase, 3-phosphoglycerate kinase, and triosephosphate isomeras," J. Bacteriol., 174(19):6076-6086 (1992)
X59404	dpg	Glutamate dehydrogenase	Bormann, E.R. et al. "Molecular analysis of the Corynebacterium glutamicum gdh gene encoding glutamate dehydrogenase," Mol. Microbiol., 6(3):317-326 (1992)
X60312	lysi	L-lysine permease	Seep-Feldhaus, A.H. et al. "Molecular analysis of the Corynebacterium glutamicum lysl gene involved in lysine uptake," Mol. Microbiol., 5(12):2995-3005 (1991)
X66078	cop1	Ps1 protein	Joliff, G. et al. "Cloning and nucleotide sequence of the csp1 gene encoding PS1, one of the two major secreted proteins of Corynebacterium glutamicum: The deduced N-terminal region of PS1 is similar to the Mycobacterium antigen 85 complex," Mol. Microbiol., 6(16):2349-2362 (1992)
X66112	glt	Citrate synthase	Eikmanns, B.J. et al. "Cloning sequence, expression and transcriptional analysis of the Corynebacterium glutamicum gltA gene encoding citrate synthase," <i>Microbiol.</i> , 140:1817-1828 (1994)
X67737	dapB	Dihydrodipicolinate reductase	
X69103	csp2	Surface layer protein PS2	Peyret, J.L. et al. "Characterization of the cspB gene encoding PS2, an ordered surface-layer protein in Corynebacterium glutamicum," Mol. Microbiol., 9(1):97-109 (1993)
X69104		IS3 related insertion element	Bonamy, C. et al. "Identification of IS1206, a Corynebacterium glutamicum IS3-related insertion sequence and phylogenetic analysis," <i>Mol. Microbiol.</i> , 14(3):571-581 (1994)

		Table 2 (continued)	nued)
X70959	leuA	Isopropylmalate synthase	Patek, M. et al. "Leucine synthesis in Corynebacterium glutamicum: enzyme activities, structure of leuA, and effect of leuA inactivation on lysine synthesis," Ann Emiron Mirrohiol, 60(1):133-140 (1994)
X71489	icd	Isocitrate dehydrogenase (NADP+)	Eikmanns, B.J. et al. "Cloning sequence analysis, expression, and inactivation of the Corynebacterium glutamicum icd gene encoding isocitrate dehydrogenase and biochemical characterization of the enzyme," J. Bacteriol., 177(3):774-782 (1995)
X72855	GDHA	Glutamate dehydrogenase (NADP+)	
X75083, X70584	mtrA	5-methyltryptophan resistance	Heery, D.M. et al. "A sequence from a tryptophan-hyperproducing strain of Corynebacterium glutamicum encoding resistance to 5-methyltryptophan," <i>Biochem. Biophys. Res. Commun.</i> , 201(3):1255-1262 (1994)
X75085	recA		Fitzpatrick, R. et al. "Construction and characterization of recA mutant strains of Corynebacterium glutamicum and Brevibacterium lactofermentum," Appl. Microbiol. Biotechnol., 42(4):575-580 (1994)
X75504	aceA; thiX	Partial Isocitrate lyase; ?	Reinscheid, D.J. et al. "Characterization of the isocitrate lyase gene from Corynebacterium glutamicum and biochemical analysis of the enzyme," J. Bacteriol., 176(12):3474-3483 (1994)
X76875		ATPase beta-subunit	Ludwig, W. et al. "Phylogenetic relationships of bacteria based on comparative sequence analysis of elongation factor Tu and ATP-synthase beta-subunit genes," Antonie Van Leeuwenhoek, 64:285-305 (1993)
X77034	tuf	Elongation factor Tu	Ludwig, W. et al. "Phylogenetic relationships of bacteria based on comparative sequence analysis of elongation factor Tu and ATP-synthase beta-subunit genes," Antonie Van Leeuwenhoek, 64:285-305 (1993)
X77384	recA		Billman-Jacobe, H. "Nucleotide sequence of a recA gene from Corynebacterium glutamicum," DNA Seq., 4(6):403-404 (1994)
X78491	aceB	Malate synthase	Reinscheid, D.J. et al. "Malate synthase from Corynebacterium glutamicum pta-ack operon encoding phosphotransacetylase: sequence analysis," <i>Microbiology</i> , 140:3099-3108 (1994)
X80629	16S rDNA	16S ribosomal RNA	Rainey, F.A. et al. "Phylogenetic analysis of the genera Rhodococcus and Norcardia and evidence for the evolutionary origin of the genus Norcardia from within the radiation of Rhodococcus species," <i>Microbiol.</i> , 141:523-528 (1995)
	gluA; gluB; gluC; gluD	Glutamate uptake system	Kronemeyer, W. et al. "Structure of the gluABCD cluster encoding the glutamate uptake system of Corynebacterium glutamicum," J. Bacteriol., 177(5):1152-1158 (1995)
X81379	dapE	Succinyldiaminopimelate desuccinylase	Wehrmann, A. et al. "Analysis of different DNA fragments of Corynebacterium glutamicum complementing dapE of Escherichia coli," <i>Microbiology</i> , 40:3349-56 (1994)

Table 2 (continued)	16S ribosomal RNA Ruimy, R. et al. "Phylogeny of the genus Corynebacterium deduced from analyses of small-subunit ribosomal DNA sequences," Int. J. Syst. Bacteriol., 45(4):740-746 (1995)	Aspartate-semialdehyde dehydrogenase; ? Serebrijski, I. et al. "Multicopy suppression by asd gene and osmotic stress-dependent complementation by heterologous proA in proA mutants," J. Bacteriol., 177(24):7255-7260 (1995)	Gamma-glutamyl phosphate reductase Serebrijski, I. et al. "Multicopy suppression by asd gene and osmotic stress-dependent complementation by heterologous proA mutants," J. Bacteriol., 177(24):7255-7260 (1995)	16S ribosomal RNA Pascual, C. et al. "Phylogenetic analysis of the genus Corynebacterium based on 16S rRNA gene sequences," <i>Int. J. Syst. Bacteriol.</i> , 45(4):724-728 (1995)	Aromatic amino acid permease; ?	Acetylglutamate kinase; N-acetyl-gamma- glutamyl-phosphate reductase; acetylornithine aminotransferase; ornithine carbamoyltransferase; glutamate N- acetyltransferase	Phosphate acetyltransferase; acetate kinase   Reinscheid, D.J. et al. "Cloning, sequence analysis, expression and inactivation of the Corynebacterium glutamicum pta-ack operon encoding   phosphotransacetylase and acetate kinase," <i>Microbiology</i> , 145:503-513 (1999)	Attachment site Le Marrec, C. et al. "Genetic characterization of site-specific integration functions of phi AAU2 infecting "Arthrobacter aureus C70," J. Bacteriol., 178(7):1996-2004 (1996)		Promoter fragment F2 Patek, M. et al. "Promoters from Corynebacterium glutamicum: cloning, molecular analysis and search for a consensus motif," <i>Microbiology</i> , 142:1297-1309 (1996)	Promoter fragment F10 Patek, M. et al. "Promoters from Corynebacterium glutamicum: cloning, molecular analysis and search for a consensus motif," Microbiology, 142:1297-1309 (1996)	Promoter fragment F13 Patek, M. et al. "Promoters from Corynebacterium glutamicum: cloning, molecular analysis and search for a consensus motif," <i>Microbiology</i> , 142:1297-1309 (1996)	
	16S ribosomal RNA	Aspartate-semialdeh	Gamma-glutamyl ph	16S ribosomal RNA		Acetylglutamate kint glutamyl-phosphate acetylornithine amin carbamoyltransferase acetyltransferase		Attachment site					
	16S rDNA	asd; lysC	proA	16S rDNA	aroP; dapE	argB; argC; argD; argF; argJ	pta; ackA	attB					
	X82061	X82928	X82929	X84257	X85965	X86157	X89084	X89850	X90356	X90357	X90358	X90359	

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ntinued)	Patek, M. et al. "Promoters from Corynebacterium glutamicum: cloning, molecular analysis and search for a consensus motif," <i>Microbiology</i> ,	Patek, M. et al. "Promoters from Corynebacterium glutamicum: cloning, molecular analysis and search for a consensus motif," <i>Microbiology</i> , 142:1297-1309 (1996)	Patek, M. et al. "Promoters from C. glutamicum: cloning, molecular analysis and search for a consensus motif," Microbiology, 142:1297-1309 (1996)	Patek, M. et al. "Promoters from Corynebacterium glutamicum: cloning, molecular analysis and search for a consensus motif," <i>Microbiology</i> , 142:1297-1309 (1996)	Patek, M. et al. "Promoters from Corynebacterium glutamicum: cloning, molecular analysis and search for a consensus motif," <i>Microbiology</i> , 142:1297-1309 (1996)	Patek, M. et al. "Promoters from Corynebacterium glutamicum: cloning, molecular analysis and search for a consensus motif," <i>Microbiology</i> , 142:1297-1309 (1996)	Patek, M. et al. "Promoters from Corynebacterium glutamicum: cloning, molecular analysis and search for a consensus motif," <i>Microbiology</i> , 142:1297-1309 (1996)	Patek, M. et al. "Promoters from Corynebacterium glutamicum: cloning, molecular analysis and search for a consensus motif," <i>Microbiology</i> , 142:1297-1309 (1996)	Patek, M. et al. "Promoters from Corynebacterium glutamicum: cloning, molecular analysis and search for a consensus motif," <i>Microbiology</i> , 142:1297-1309 (1996)	Siewe, R.M. et al. "Functional and genetic characterization of the (methyl) ammonium uptake carrier of Corynebacterium glutamicum," J. Biol. Chem., 271(10):5398-5403 (1996)	Peter, H. et al. "Isolation, characterization, and expression of the Corynebacterium glutamicum betP gene, encoding the transport system for the compatible solute glycine betaine," J. Bacteriol., 178(17):5229-5234 (1996)	Patek, M. et al. "Identification and transcriptional analysis of the dapB-ORF2-dapA-ORF4 operon of Corynebacterium glutamicum, encoding two enzymes involved in L-lysine synthesis," Biotechnol. Lett., 19:1113-1117 (1997)	Vrljic, M. et al. "A new type of transporter with a new type of cellular function: L-lysine export from Corynebacterium glutamicum," Mol. Microbiol., 22(5):815-826 (1996)
Table 2 (continued)	Promoter fragment F22	Promoter fragment F34	Promoter fragment F37	Promoter fragment F45	Promoter fragment F64	Promoter fragment F75	Promoter fragment PF101	Promoter fragment PF104	Promoter fragment PF109	Ammonium transport system	Glycine betaine transport system		Lysine exporter protein; Lysine export regulator protein
										amt	betP	orf4	lysE; lysG
	X90360	X90361	X90362	X90363	X90364	X90365	X90366	X90367	X90368	X93513	X93514	X95649	X96471

		Table 2 (continued)	(penu
X96580	panB; panC; xylB	3-methyl-2-oxobutanoate hydroxymethyltransferase; pantoate-beta- alanine ligase; xylulokinase	Sahm, H. et al. "D-pantothenate synthesis in Corynebacterium glutamicum and use of panBC and genes encoding L-valine synthesis for D-pantothenate overproduction," <i>Appl. Environ. Microbiol.</i> , 65(5):1973-1979 (1999)
X96962		Insertion sequence IS1207 and transposase	
X99289		Elongation factor P	Ramos, A. et al. "Cloning, sequencing and expression of the gene encoding elongation factor P in the amino-acid producer Brevibacterium lactofermentum (Corynebacterium glutamicum ATCC 13869)," Gene, 198:217-222 (1997)
Y00140	thrB	Homoserine kinase	Mateos, L.M. et al. "Nucleotide sequence of the homoserine kinase (thrB) gene of the Brevibacterium lactofermentum," <i>Nucleic Acids Res.</i> , 15(9):3922 (1987)
Y00151	qpp	Meso-diaminopimelate D-dehydrogenase (EC 1.4.1.16)	Ishino, S. et al. "Nucleotide sequence of the meso-diaminopimelate D-dehydrogenase gene from Corynebacterium glutamicum," Nucleic Acids Res., 15(9):3917 (1987)
Y00476	thrA	Homoserine dehydrogenase	Mateos, L.M. et al. "Nucleotide sequence of the homoserine dehydrogenase (thrA) gene of the Brevibacterium lactofermentum," <i>Nucleic Acids Res.</i> , 15(24):10598 (1987)
Y00546	hom; thrB	Homoserine dehydrogenase; homoserine kinase	Peoples, O.P. et al. "Nucleotide sequence and fine structural analysis of the Corynebacterium glutamicum hom-thrB operon," Mol. Microbiol., 2(1):63-72 (1988)
Y08964	murC; ftsQ/divD; ftsZ	UPD-N-acetylmuramate-alanine ligase; division initiation protein or cell division protein; cell division protein	Honrubia, M.P. et al. "Identification, characterization, and chromosomal organization of the ftsZ gene from Brevibacterium lactofermentum," Mol. Gen. Genet., 259(1):97-104 (1998)
Y09163	putP	High affinity proline transport system	Peter, H. et al. "Isolation of the putP gene of Corynebacterium glutamicumproline and characterization of a low-affinity uptake system for compatible solutes," <i>Arch. Microbiol.</i> , 168(2):143-151 (1997)
Y09548	pyc	Pyruvate carboxylase	Peters-Wendisch, P.G. et al. "Pyruvate carboxylase from Corynebacterium glutamicum: characterization, expression and inactivation of the pyc gene," <i>Microbiology</i> , 144:915-927 (1998)
Y09578	leuB	3-isopropylmalate dehydrogenase	Patek, M. et al. "Analysis of the leuB gene from Corynebacterium glutamicum," Appl. Microbiol. Biotechnol., 50(1):42-47 (1998)
Y12472		Attachment site bacteriophage Phi-16	Moreau, S. et al. "Site-specific integration of corynephage Phi-16: The construction of an integration vector," <i>Microbiol.</i> , 145:539-548 (1999)
Y12537	proP	Proline/ectoine uptake system protein	Peter, H. et al. "Corynebacterium glutamicum is equipped with four secondary carriers for compatible solutes: Identification, sequencing, and characterization of the proline/ectoine uptake system, ProP, and the ectoine/proline/glycine betaine carrier, EctP," J. Bacteriol., 180(22):6005-6012 (1998)

		Table 2 (continued)	ned)
Y13221	glnA	Glutamine synthetase I	Jakoby, M. et al. "Isolation of Corynebacterium glutamicum glnA gene encoding glutamine synthetase I," FEMS Microbiol. Lett., 154(1):81-88 (1997)
Y16642	lpd!	Dihydrolipoamide dehydrogenase	
Y18059		Attachment site Corynephage 304L	Moreau, S. et al. "Analysis of the integration functions of φ304L: An integrase module among corynephages," Virology, 255(1):150-159 (1999)
221501	argS; lysA	Arginyl-tRNA synthetase; diaminopimelate decarboxylase (partial)	Oguiza, J.A. et al. "A gene encoding arginyl-tRNA synthetase is located in the upstream region of the lysA gene in Brevibacterium lactofermentum: Regulation of argS-lysA cluster expression by arginine," J. Bacteriol., 175(22):7356-7362 (1993)
Z21502	dapA; dapB	Dihydrodipicolinate synthase; dihydrodipicolinate reductase	Pisabarro, A. et al. "A cluster of three genes (dapA, orf2, and dapB) of Brevibacterium lactofermentum encodes dihydrodipicolinate reductase, and a third polypeptide of unknown function," J. Bacteriol., 175(9):2743-2749 (1993)
Z29563	thrC	Threonine synthase	Malumbres, M. et al. "Analysis and expression of the thrC gene of the encoded threonine synthase," Appl. Environ. Microbiol., 60(7)2209-2219 (1994)
Z46753	16S rDNA	Gene for 16S ribosomal RNA	
Z49822	sigA	SigA sigma factor	Oguiza, J.A. et al "Multiple sigma factor genes in Brevibacterium lactofermentum: Characterization of sigA and sigB," J. Bacteriol., 178(2):550-553 (1996)
Z49823	galE; dtxR	Catalytic activity UDP-galactose 4- epimerase; diphtheria toxin regulatory protein	Oguiza, J.A. et al "The galE gene encoding the UDP-galactose 4-epimerase of Brevibacterium lactofermentum is coupled transcriptionally to the dmdR gene," Gene, 177:103-107 (1996)
Z49824	orfl; sigB	?; SigB sigma factor	Oguiza, J.A. et al "Multiple sigma factor genes in Brevibacterium lactofermentum: Characterization of sigA and sigB," J. Bacteriol., 178(2):550-553 (1996)
Z66534		Transposase	Correia, A. et al. "Cloning and characterization of an IS-like element present in the genome of Brevibacterium lactofermentum ATCC 13869," <i>Gene</i> , 170(1):91-94 (1996)
A sequence for the published ve.	' A sequence for this gene was published in the indicat the published version. It is believed that the published	the indicated reference. However, the sequence published version relied on an incorrect start c	ted reference. However, the sequence obtained by the inventors of the present application is significantly longer than version relied on an incorrect start codon, and thus represents only a fragment of the actual coding region.

TABLE 3: Corynebacterium and Brevibacterium Strains Which May be Used in the Practice of the Invention

Genus # 25	species 🔆 📜 🚡	ATCC	FERM	NRRE	CECT	NCIMB	<b>CBS</b>	NCTE	DSMZ
Brevibacterium	ammoniagenes	21054							
Brevibacterium	ammoniagenes	19350				-			
Brevibacterium	ammoniagenes	19351							
Brevibacterium	ammoniagenes	19352							
Brevibacterium	ammoniagenes	19353							
Brevibacterium	ammoniagenes	19354							
Brevibacterium	ammoniagenes	19355							
Brevibacterium	ammoniagenes	19356							
Brevibacterium	ammoniagenes	21055							
Brevibacterium	ammoniagenes	21077							
Brevibacterium	ammoniagenes	21553							
Brevibacterium	ammoniagenes	21580							
Brevibacterium	ammoniagenes	39101							
Brevibacterium	butanicum	21196							
Brevibacterium	divaricatum	21792	P928						
Brevibacterium	flavum	21474							
Brevibacterium	flavum	21129							
Brevibacterium	flavum	21518							
Brevibacterium	flavum			B11474					
Brevibacterium	flavum			B11472					
Brevibacterium	flavum	21127							
Brevibacterium	flavum	21128							
Brevibacterium	flavum	21427							
Brevibacterium	flavum	21475							
Brevibacterium	flavum	21517							
Brevibacterium	flavum	21528							
Brevibacterium	flavum	21529							
Brevibacterium	flavum			B11477	,,,				
Brevibacterium	flavum			B11478					
Brevibacterium	flavum	21127							
Brevibacterium	flavum			B11474					
Brevibacterium	healii	15527							
Brevibacterium	ketoglutamicum	21004							
Brevibacterium	ketoglutamicum	21089							
Brevibacterium	ketosoreductum	21914							
Brevibacterium	lactofermentum				70				
Brevibacterium	lactofermentum				74				
Brevibacterium	lactofermentum				77				
Brevibacterium	lactofermentum	21798							
Brevibacterium	lactofermentum	21799							
Brevibacterium	lactofermentum	21800							
Brevibacterium	lactofermentum	21801							
Brevibacterium	lactofermentum			B11470					
Brevibacterium	lactofermentum			B11471					

	species		FERM	NRRL	CECT	NCIMB	CBS	NCTC	DSMZ
Brevibacterium	lactofermentum	21086							
Brevibacterium	lactofermentum	21420							
Brevibacterium	lactofermentum	21086							
Brevibacterium	lactofermentum	31269							
Brevibacterium	linens	9174							
Brevibacterium	linens	19391							
Brevibacterium	linens	8377							
Brevibacterium	paraffinolyticum					11160			
Brevibacterium	spec.						717.73		
Brevibacterium	spec.		<u> </u>				717.73		
Brevibacterium	spec.	14604							
Brevibacterium	spec.	21860							
Brevibacterium	spec.	21864	1						
Brevibacterium	spec.	21865							
Brevibacterium	spec.	21866							
Brevibacterium	spec.	19240	1						
Corynebacterium	acetoacidophilum	21476							
Corynebacterium	acetoacidophilum	13870	<u> </u>						
Corynebacterium	acetoglutamicum			B11473					
Corynebacterium	acetoglutamicum			B11475					
Corynebacterium	acetoglutamicum	15806							
Corynebacterium	acetoglutamicum	21491							
Corynebacterium	acetoglutamicum	31270	1						
Corynebacterium	acetophilum		1	B3671					
Corynebacterium	ammoniagenes	6872	1					2399	
Corynebacterium	ammoniagenes	15511	1					l	
Corynebacterium	fujiokense	21496							
Corynebacterium	glutamicum	14067							
Corynebacterium	glutamicum	39137	T						
Corynebacterium	glutamicum	21254							
Corynebacterium	glutamicum	21255							
Corynebacterium	glutamicum	31830							
Corynebacterium	glutamicum	13032			-				
Corynebacterium	glutamicum	14305							
Corynebacterium	glutamicum	15455							
Corynebacterium	glutamicum	13058							
Corynebacterium	glutamicum	13059							
Corynebacterium	glutamicum	13060							
Corynebacterium	glutamicum	21492							
Corynebacterium	glutamicum	21513							
Corynebacterium	glutamicum	21526							
Corynebacterium	glutamicum	21543							
Corynebacterium	glutamicum	13287							
Corynebacterium	glutamicum	21851							
Corynebacterium	glutamicum	21253							
Corynebacterium	glutamicum	21514							
Corynebacterium	glutamicum	21516	T						
Corynebacterium	glutamicum	21299	1						

Corynebacterium   Corynebact	Genus 🔭 🦠	species 📜 🖫	ATCC	FERM	NRRL	CECT	NGIMB	CBS	NCTC	DSMZ
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Corynebacterium   glutamicum   21488		<u> </u>	39684							
Corynebacterium   glutamicum   21649		glutamicum	21488			<b>T</b>				
Corynebacterium   Corynebact			21649							
Corynebacterium   glutamicum   1923		glutamicum	21650			<u> </u>				
Corynebacterium   glutamicum   21157		glutamicum	19223							
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Corynebacterium   glutamicum   21568	Corynebacterium	glutamicum	21567							
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Corynebacterium         glutamicum         21573           Corynebacterium         glutamicum         21579           Corynebacterium         glutamicum         19049           Corynebacterium         glutamicum         19050           Corynebacterium         glutamicum         19051           Corynebacterium         glutamicum         19052           Corynebacterium         glutamicum         19053           Corynebacterium         glutamicum         19054           Corynebacterium         glutamicum         19055           Corynebacterium         glutamicum         19056           Corynebacterium         glutamicum         19058           Corynebacterium         glutamicum         19059           Corynebacterium         glutamicum         19185           Corynebacterium         glutamicum         19185           Corynebacterium         glutamicum         21515           Corynebacterium         glutamicum         21527           Corynebacterium         glutamicum         21544           Corynebacterium         glutamicum         21492           Corynebacterium         glutamicum         21492           Corynebacterium         glutamicum         21492	Corynebacterium	glutamicum	21571							
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[ , _ ·	Corynebacterium	glutamicum			B8182					
Corynebacterium glutamicum B12416		T			B12416	<u> </u>				
Corynebacterium glutamicum B12417	Corynebacterium	glutamicum			B12417					

Genus	species	#ATCC	FERM	NRRL	CECT	NCIMB	CBS	NCTC	DSMZ
Corynebacterium	glutamicum			B12418	[ _				
Corynebacterium	glutamicum			B11476					
Corynebacterium	glutamicum	21608							
Corynebacterium	lilium		P973						
Corynebacterium	nitrilophilus	21419				11594			
Corynebacterium	spec.		P4445						
Corynebacterium	spec.		P4446						
Corynebacterium	spec.	31088							
Corynebacterium	spec.	31089							
Corynebacterium	spec.	31090							
Corynebacterium	spec.	31090							
Corynebacterium	spec.	31090							
Corynebacterium	spec.	15954							20145
Corynebacterium	spec.	21857							
Corynebacterium	spec.	21862							
Corynebacterium	spec.	21863							

ATCC: American Type Culture Collection, Rockville, MD, USA

FERM: Fermentation Research Institute, Chiba, Japan

NRRL: ARS Culture Collection, Northern Regional Research Laboratory, Peoria, IL, USA

CECT: Coleccion Espanola de Cultivos Tipo, Valencia, Spain

NCIMB: National Collection of Industrial and Marine Bacteria Ltd., Aberdeen, UK

CBS: Centraalbureau voor Schimmelcultures, Baarn, NL

NCTC: National Collection of Type Cultures, London, UK

DSMZ: Deutsche Sammlung von Mikroorganismen und Zellkulturen, Braunschweig, Germany

For reference see Sugawara, H. et al. (1993) World directory of collections of cultures of microorganisms: Bacteria, fungi and yeasts (4<sup>th</sup> edn), World federation for culture collections world data center on microorganisms, Saimata, Japen.

				Table 4: Alignment Results			
ID# length	length Genbank Hit (NT)	Length	Length Accession Na	me of Genbank	Source of Genbank Hit	% homology, Date of (GAP)	Date of Deposit
rxa00013 996	GB_GSS4:AQ713475	581	AQ713475	HS_5402_B2_A12_T7A RPCI-11 Human Male BAC Library Homo sapiens	Homo sapiens	37,148	13-Jul-99
	GB_HTG3:AC007420	130583	130583 AC007420	9cronnic done in the control of the	Drosophila melanogaster	34,568	20-Sep-99
	GB_HTG3:AC007420	130583	130583 AC007420	, 83 unordered pieces. osophila melanogaster chromosome 2 clone BACR07M10 (D630) RPCI-98 .M.10 map 24A-24D strain y; cn bw sp, *** SEQUENCING IN :OGRESS***, 83 unordered pieces.	Drosophila melanogaster	34,568	20-Sep-99
rxa00014 903	GB_BA1:MTCY3A2	25830	Z83867	Mycobacterium tuberculosis H37Rv complete genome; segment 136/162.	Mycobacterium	58,140	17-Jun-98
	GB_BA1:MLCB1779		Z98271	Mycobacterium leprae cosmid B1779.	Mycobacterium leprae	57,589	8-Aug-97
643 06000	GB_BA1:SAPURCLUS	9120	X92429	S.alboniger napH, pur7, pur10, pur6, pur4, pur5 and pur3 genes.  Steptomyces anulatus C80712 Picture disoridation of the pure disoridation disoridation disoridation disoridation disoridation disoridation.	Streptomyces anulatus	55,667	28-Feb-96
000000000000000000000000000000000000000	CD		2 200	CONTINUISM INTERPRETATION OF CONTINUISM IN DICTOR CONTINUISM CONTINUISM IN CONTINUISM INCOME.	Orciyostenam discolaeum	45,265	66-1d4-02
	GB_EST28:AI497294	<b>4</b> 8	A1497294	fb63g03.y1 Zebrafish WashU MPIMG EST Danio rerio cDNA 5' similar to SW:AFP4_MYOOC P80961 ANTIFREEZE PROTEIN LS-12: ;; mRNA sequence.	Danio rerio	42,991	11-MAR-1999
	GB_EST21:C92167	637	C92167	C92167 Dictyostelium discoideum SS (H. Urushihara) Dictyostelium discoideum Dictyostelium discoideum cDNA clone SSD179, mRNA sequence.	Dictyostelium discoideum	44,444	12-Jul-99
xa00032 1632	GB_BA2:AF010496	189370	189370 AF010496	Rhodobacter capsulatus strain SB1003, partial genome.	Rhodobacter capsulatus	39,689	12-MAY-1998
	GB_BA2:AF018073	9810	AF018073	Rhodobacter sphaeroides operon regulator (smoC), periplasmic sorbitol-binding Rhodobacter sphaeroides protein (smoE), sorbitol/mannitol transport inner membrane protein (smoF), sorbitol/mannitol transport inner membrane protein (smoG), sorbitol/mannitol transport protein (smoK), sorbitol dehydrogenase (smoS), mannitol dehydrogenase (mtlK), and periplasmic mannitol-binding protein (smoM) genes, complete cds.	Rhodobacter sphaeroides	48,045	22-OCT-1997
	GB_BA2:AF045245	5930	AF045245	Klebsiella pneumoniae D-arabinitol transporter (daIT), D-arabinitol kinase (daIK), D-arabinitol dehydrogenase (daID), and repressor (daIR) genes, complete cds.	Klebsiella pneumoniae	38,514	16-Jul-98
rxa000041 1342	EM_PAT:E11760	6911	E11760	Base sequence of sucrase gene.	Corynebacterium glutamicum	99,031	08-OCT-1997 (Rel. 52, Created)
	GB_PAT:126124	6911	126124	Sequence 4 from patent US 5556776.	Unknown.	99,031	07-OCT-1996
	GB_IN1:LMFL5883	31934	AL117384		Leishmania major	43,663	21-OCT-1999
xa00042 882	EM_PAT:E11760	6911	E11760	Base sequence of sucrase gene.	Corynebacterium glutamicum	94,767	08-OCT-1997 (Rel. 52, Created)
	GB_PAT:126124	6911	126124	Sequence 4 from patent US 5556776.	Unknown.	94,767	07-OCT-1996

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23-Jan-96 07-OCT-1996 08-OCT-1997 (Rel. 52,	24-Jun-98 19-Apr-97	27-Jul-98 17-Jun-98	17-Jun-98	03-DEC-1996	03-DEC-1996	10-DEC-1996	17-Jun-98	19-Jun-98	15-Jun-96 19-Jun-98	19-Jun-98	19-Jun-98	27-Apr-93 13-MAR-1997	2-Feb-99	2-Feb-99	17-Sep-97 23-Jun-99
40,276 97,591 97,591	35,879 s 62,658	37,638 36,784	67,457	40,883	35,883	51,001	51,001	35,735	57,014 41,892	41,841	36,599	36,212 38,816	42,239	37,307	58,312 36,632
Caenorhabditis elegans Unknown. Corynebacterium glutamicum	Homo sapiens 35,879 Mycobacterium smegmatis 62,658	Streptomyces coelicolor Mycobacterium	Mycobacterium tuberculosis	Mycobacterium tuberculosis	tuberculosis Mycobacterium	tuberculosis Mycobacterium	Mycobacterium	ruberculosis Mycobacterium	Mycobacterium leprae Mycobacterium	tuberculosis Mycobacterium tuberculosis	Mycobacterium	Rattus norvegicus	5' Mus musculus	Mus musculus	Mycobacterium leprae Mycobacterium tuberculosis
Table 4 (continued) Caenorhabditis elegans sur-2 mRNA, complete cds. Sequence 4 from patent US 5556776. Base sequence of sucrase gene.	Homo sapiens clone UWGC:g1564a012 from 7p14-15, complete sequence. Mycobacterium smegmatis phosphoglucose isomerase gene, complete cds.	Streptomyces coelicolor cosmid 5A7. Mycobacterium tuberculosis H37Rv complete genome; segment 44/162.	Mycobacterium tuberculosis H37Rv complete genome; segment 65/162.	AD000001 Mycobacterium tuberculosis sequence from clone y456.  AD000015 Mycobacterium tuberculosis sequence from clone y456.		Mycobacterium tuberculosis sequence from clone y175.	Mycobacterium tuberculosis H37Rv complete genome; segment 65/162.	Mycobacterium tuberculosis H37Rv complete genome; segment 126/162.	Mycobacterium leprae cosmid B1529 DNA sequence. Mycobacterium tuberculosis H37Rv complete genome; segment 126/162.	Mycobacterium tuberculosis H37Rv complete genome; segment 126/162.	Mycobacterium tuberculosis H37Rv complete genome; segment 126/162.	Rat carbohydrate binding receptor gene, complete cds. mw95c10.r1 Soares mouse NML Mus musculus cDNA clone IMAGE:678450 5', Mus musculus	mRNA sequence. mw96a03.y1 Soares mouse NML Mus musculus cDNA clone IMAGE:678508 5' Mus musculus similar to TR:009171 009171 BETAINE-HOMOCYSTEINE METHYLTRANSFERASE: mRNA sequence.	mw95c10.y1 Soares mouse NML Mus musculus cDNA clone IMAGE:678450 5' mRNA sequence	Mycobacterium leprae cosmid B637.  Mycobacterium tuberculosis H37Rv complete genome; segment 132/162.
U33051 126124 E11760	AC005174 U88433	AL031107 Z79700	Z79701	AD000001	AD000001	AD000015	279701	274024	L78824 Z74024	Z74024	Z74024	M55532 AA253618	AI390284	AI390280	299263 AL021287
4899 6911 6911	39769 1928	40337 39800	38300	37316	37316	18106	38300	39991	36985 39991	39991	39991	10752 313	490	467	44882 70287
GB_IN1:CEU33051 GB_PAT:I26124 EM_PAT:E11760	GB_PR3:AC005174 GB_BA1:MSU88433	GB_BA1:SC5A7 GB_BA1:MTCY10D7	GB_BA1:MTCY277	GB_BA1:MSGY456	GB_BA1:MSGY456	GB_BA1:MSGY175	GB_BA1:MTCY277	GB_BA1:MTCY274	GB_BA1:MSGB1529CS 36985 GB_BA1:MTCY274 39991	GB_BA1:MTCY274	GB_BA1:MTCY274	GB_RO:RATCBRQ GB_EST11:AA253618	GB_EST26:Al390284	GB_EST26:Al390280	GB_BA1:MLCB637 GB_BA1:MTV012
ка00043 1287	rxa00098 1743		rxa00148 2334		rxa00149 1971			rxa00195 684		rxa00196 738		xa00202 1065			xa00206 1161

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				Table 4 (continued)			
	GB_PR1:HUMFMO1	2134	M64082	Js.	Homo sapiens	37,915	8-Nov-94
	GB_EST32:AI734238	512	AI734238	zb73c05.y5 Soares_fetal_lung_NbHL19W Homo sapiens cDNA clone IMAGE:309224 5' similar to gb:M64082 DIMETHYLANILINE MONOOXYGENASE (HUMAN);, mRNA sequence.	Homo sapiens	41,502	14-Jun-99
гха00296 2967	GB_HTG6:AC011069	168266	168266 AC011069	CR11H20 (D881) RPCI-98 JENGING IN PROGRESS	Drosophila melanogaster	33,890	02-DEC-1999
	GB_EST15:AA531468	414	AA531468	Pr10 Homo sapiens cDNA clone IMAGE:997175,	Homo sapiens	40,821	20-Aug-97
	GB_HTG6:AC011069	168266	168266 AC011069	ogaster chromosome X clone BACR11H20 (D881) RPCI-98 -12C strain y; cn bw sp, *** SEQUENCING IN PROGRESS pieces.	Drosophila melanogaster	30,963	02-DEC-1999
rxa00310 558	GB_VI:VMVY16780	186986	Y16780	variola minor virus complete genome.	variola minor virus	35,883	2-Sep-99
	GB_VI:VARCG	186103		1-1975) complete genome.	Variola major virus	34,664	12-Jan-95
	GB_VI:VVCGAA	185578			Variola virus	36,000	13-DEC-1996
гха00317 777	GB_HTG3:AC009571	159648	AC009571	Homo sapiens chromosome 4 clone 57_A_22 map 4, *** SEQUENCING IN PROGRESS ***, 8 unordered pieces.	Homo sapiens	36,988	29-Sep-99
	GB_HTG3:AC009571	159648	AC009571	Homo sapiens chromosome 4 clone 57_A_22 map 4, *** SEQUENCING IN PROGRESS ***, 8 unordered pieces.	Homo sapiens	36,988	29-Sep-99
	GB_PR3:AC005697	174503	AC005697	Homo sapiens chromosome 17, clone hRPK.138_P_22, complete sequence.	Homo sapiens	36,340	09-OCT-1998
rxa00327 507	GB_BA1:LCATPASEB	1514	X64542	L.casei gene for ATPase beta-subunit.	Lactobacillus casei	34,664	11-DEC-1992
	GB_BA1:LCATPASEB	1514	X64542	L. casei gene for ATPase beta-subunit.	Lactobacillus casei	39,308	11-DEC-1992
ка00328 615	GB_BA1:STYPUTPE	1887	L01138	Salmonella (S2980) proline permease (putP) gene, 5' end.	Salmonella sp.	39,623	09-MAY-1996
	GB_BA1:STYPUTPF	1887	L01139	Salmonella (S2983) proline permease (putP) gene, 5' end.	Salmonella sp.	39,623	09-MAY-1996
	GB_BA1:STYPUTPI	1889	L01142	Salmonella (S3015) proline permease (putP) gene, 5' end.	Salmonella sp.	42,906	09-MAY-1996
rxa00329 1347	GB_PR3:AC004691	141990	AC004691	Homo sapiens PAC clone DJ0740D02 from 7p14-p15, complete sequence.	Homo sapiens	38,142	16-MAY-1998
	GB_PR4:AC004916 GB_PR3:AC004691	129014 141990	129014 AC004916 141990 AC004691	Homo sapiens clone DJ0891L14, complete sequence. Homo sapiens PAC clone DJ0740D02 from 7p14-p15, complete sequence.	Homo sapiens Homo sapiens	38,549 35,865	17-Jul-99 16-MAY-1998
ка00340 1269	GB_BA1:MTCY427	38110	270692	Mycobacterium tuberculosis H37Rv complete genome; segment 99/162.	Mycobacterium tuberculosis	38,940	24-Jun-99
	GB_GSS12:AQ412290	238	AQ412290	RPCI-11-195H2.TV RPCI-11 Homo sapiens genomic clone RPCI-11-195H2,	Homo sapiens	36,555	23-MAR-1999
	GB_PL2:AF112871	2394	AF112871	genomic solvey sequence. Astasia longa small subunit ribosomal RNA gene, complete sequence.	Astasia longa	36,465	28-Jun-99
rxa00379 307	GB_HTG1:CEY56A3	224746	AL022280	Caenorhabditis elegans chromosome III clone Y56A3, *** SEQUENCING IN PROGRESS ***, in unordered pieces.	Caenorhabditis elegans	35,179	6-Sep-99
	GB_HTG1:CEY56A3	224746	224746 AL022280	Caenorhabditis elegans chromosome III clone Y56A3, *** SEQUENCING IN PROGRESS ***, in unordered pieces.	Caenorhabditis elegans	35,179	6-Sep-99

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23-Nov-99	15-Jul-99	2-Sep-98	2-Sep-98	11-Jun-99	16-Jul-99	29-MAR-1996	17-Jun-98	03-DEC-1996	13-Sep-99	10-DEC-1996	17-Jun-98	29-MAY-1999	22-Aug-97	01-MAR-1999	23-Feb-99	23-Nov-99	23-Nov-99	17-Jun-98
40,604	35,766	41,113	41,113	41,152	41,360	36,792	51,852	51,852	36,875	60,022	60,022	28,013	38,226	37,492	36,648	39,831	36,409	56,232
Homo sapiens	Homo sapiens	e Mus musculus	e Mus musculus	Gossypium hirsutum	Homo sapiens	Homo sapiens	Mycobacterium tuberculosis	Mycobacterium tuberculosis	Homo sapiens	Mycobacterium tuberculosis	Mycobacterium tuberculosis	Caenorhabditis elegans	Homo sapiens	Lactococcus lactis subsp.	Caenorhabditis elegans	Homo sapiens	Homo sapiens	Mycobacterium tuberculosis
Human DNA sequence from clone 134019 on chromosome 1p36.11-36.33,	를 유 등 등 등 등 등 등 등 등 등 등 등 등 등 등 등 등 등 등	ub74f05.r1 Soares mouse mammary gland NMLMG Mus musculus cDNA clone Mus musculus IMAGE:1383489 5' similar to gb:J04046 CALMODULIN (HUMAN); gb:M19381 Mouse calmodulin (MOUSE);, mRNA sequence.	ub74f05.r1 Soares mouse mammary gland NMLMG Mus musculus cDNA clone Mus musculus IMAGE:1383489 5' similar to gb:J04046 CALMODULIN (HUMAN); gb:M19381 Mouse calmodulin (MOUSE);, mRNA sequence.	BNLGHi5857 Six-day Cotton fiber Gossypium hirsutum cDNA 5' similar to (AF015913) Skb1Hs [Homo sapiens], mRNA sequence.		H.sapiens mRNA for GAIP protein.	Mycobacterium tuberculosis H37Rv complete genome; segment 28/162.		Homo sapiens chromosome 21 clone B2308H15 map 21q22.3, *** SEQUENCING IN PROGRESS ***, in unordered pieces.	Mycobacterium tuberculosis sequence from clone y126.	Mycobacterium tuberculosis H37Rv complete genome; segment 156/162.	Caenorhabditis elegans chromosome II clone Y48C3, *** SEQUENCING IN PROGRESS ***, in unordered pieces.	Homo sapiens chromosome 16 BAC clone CIT987SK-334D11 complete sequence.	Lactococcus lactis cremoris plasmid pJW565 DNA, abiiM, abiiR genes and orfX.		Human DNA sequence from cosmid E127C11 on chromosome 22q11.2-qter contains STS.	Human DNA sequence from cosmid E127C11 on chromosome 22q11.2-qter contains STS.	Mycobacterium tuberculosis H37Rv complete genome; segment 49/162.
AL034555	AQ730532	AI120939	AI120939	AI726450	AQ740856	X91809	295558	AD000004	AP000471	AD000012	Z80343	270193 292855	AF001550	Y12736	206217 AC006754	274581	<b>Z</b> 74581	Z95585
86897	416	261	561	565	768	1587	40838	40051	72466	37164	37085	270193	173882	12828	206217	38423	38423	22550
GB_PR2:HS134019	GB_GSS4:AQ730532	GB_EST23:Al120939	GB_EST23:AI120939	GB_EST32:AI726450	GB_GSS4:AQ740856	GB_PR1:HSPAIP	GB_BA1:MTY25D10	GB_BA1:MSGY224	GB_HTG1:AP000471	GB_BA1:MSGY126	GB_BA1:MTY13D12	GB_HTG1:CEY48C3	GB_PR2:HSAF001550 173882 AF001550	GB_BA1:LCPJW565	GB_HTG2:AC006754	GB_PR3:HSE127C11	GB_PR3:HSE127C11	GB_BA1:MTCY22G8
	гха00381 729			rxa00385 362			rxa00388 1134			rxa00427 909			rxa00483 1587			гха00511 615		xa00512 718

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				Table 4 (continued)			
rxa00683 1215	GB_BA2:AE000896	10707	AE000896	Methanobacterium thermoautotrophicum from bases 1189349 to 1200055	Methanobacterium	38,429	15-Nov-97
				(section 102 of 148) of the complete genome.	thermoautotrophicum		
	GB_IN1:DMBR7A4	212734		Drosophila melanogaster clone BACR7A4.	Drosophila melanogaster	36,454	30-Jul-99
	GB_EST35:AV163010	273	AV163010	AV163010 Mus musculus head C57BL/6J 13-day embryo Mus musculus cDNA Mus musculus	. Mus musculus	41,758	8-Jul-99
				clone 3110006J22, mRNA sequence.			
rxa00686 927	GB_HTG2:HSDJ137K2 190223 AL049820	190223	AL049820	Homo sapiens chromosome 6 clone RP1-137K2 map q25.1-25.3, ***	Homo sapiens	38,031	03-DEC-1999
	GB_HTG2:HSDJ137K2 190223 AL049820	190223	AL049820	Homo sapiens chromosome 6 clone RP1-137K2 map q25.1-25.3, ***	Homo sapiens	38,031	03-DEC-1999
				SEQUENCING IN PROGRESS ***, in unordered pieces.			
	GB_EST12:AA284399	431	AA284399	zs57b04.r1 NCI_CGAP_GCB1 Homo sapiens cDNA clone IMAGE:701551 5;	Homo sapiens	39,205	14-Aug-97
			٠	mRNA sequence.			
rxa00700 927	GB_EST34:AI785570	45 4	AI785570	uj44d03.x1 Sugano mouse liver mlia Mus musculus cDNA clone	Mus musculus	41,943	2-Jul-99
				IMAGE: 1922/89 3' SIMIIAT to gb: 22640/ 605 RIBOSOMAL PROTEIN L8			
	CD ECTOR: A1266147	700	A1256147	(NOMAN), FINANA Sequence.	Mus museulus	40 704	12 Mov.08
	110071007107-00	ţ	21.0031	IMAGE:1890190 3' similar to ob: 728407 60S RIBOSOMAL PROTEIN 18			00-4041-71
				(HUMAN); mRNA sequence.			
	GB_BA1:CARCG12	2079	X14979	C. aurantiacus reaction center genes 1 and 2.	Chloroflexus aurantiacus	37,721	23-Apr-91
rxa00703 2409	GB_BA1:SC7H2	42655	AL109732	Streptomyces coelicolor cosmid 7H2.	Streptomyces coelicotor	56,646	2-Aug-99
	1				A3(2)		,
	GB_BA1:MTCY274	39991	Z74024	Mycobacterium tuberculosis H37Rv complete genome; segment 126/162.	Mycobacterium	37,369	19-Jun-98
					tuperculosis		
	GB_BA2:REU60056	2520	N60056	Raistonia eutropha formate dehydrogenase-like protein (cbbBc) gene, complete Raistonia eutropha cds.	e Ralstonia eutropha	51,087	16-OCT-1996
rxa00705 1038	GB_GSS15.AQ604477	505	AQ604477	HS_2116_B1_G07_MR CIT Approved Human Genomic Sperm Library D Homo Homo sapiens sapiens genomic clone Plate=2116 Col=13 Row=N, genomic survey sequence.	o Homo sapiens	39,617	10-Jun-99
	0.000 de 2.1140 T		4			6	
	GB_ES111:AA224340	443	AACZ4340	ZT146U7.S1 Stratagene nN I neuron (#33/233) nomo sapiens cunva cione IMAGE:648804 3', mRNA sequence.	nomo sapiens	35,129	11-WAK-1998
	GB_EST5:N30648	291	N30648	yw77b02.s1 Soares_placenta_8to9weeks_2NbHP8to9W Homo sapiens cDNA	Homo sapiens	43,986	5-Jan-96
rxa00782 1005	GB BA1:MTCY10D7	39800	279700	Mycobacterium tuberculosis H37Rv complete genome; segment 44/162.	Mycobacterium	63,327	17-Jun-98
	1				tuberculosis		
	GB_BA1:MLCL373	37304	AL035500	Mycobacterium leprae cosmid L373.	Mycobacterium leprae		27-Aug-99
	GB_BA2:AF128399	2842	AF128399	Pseudomonas aeruginosa succinyi-CoA synthetase beta subunit (sucC) and succinyi-CoA synthetase alpha subunit (sucD) genes complete cds	Pseudomonas aeruginosa	53,698	25-MAR-1999
rxa00783 1395	GB_HTG2:AC008158	118792	118792 AC008158		Homo sapiens	35,135	28-Jul-99
	GB_HTG2:AC008158	118792	118792 AC008158	Homo sapiens chromosome 17 clone hRPK.42_F_20 map 17, *** SEOI IFNCING IN PROCRESS *** 14 unordered nieces	Homo sapiens	35,135	28-Jul-99
	GB PR3:AC005017	137176	137176 AC005017		Homo sapiens	35,864	8-Aug-98
rxa00794 1128	GB_BA1:MTV017	67200	AL021897		Mycobacterium	40,331	24-Jun-99
					tuberculosis		

				Table 4 (continued)			
	GB_BA1:MLCB1222 GB_PR2:HS151B14	34714	34714 AL049491 128942 Z82188	Mycobacterium leprae cosmid B1222.  Human DNA sequence from clone 151B14 on chromosome 22 Contains SOMATOSTATIN RECEPTOR TYPE 3 (SS3R) gene, pseudogene similar to ribosomal protein L39,RAC2 (RAS-RELATED C3 BOTULINUM TOXIN SUBSTRATE (P21-RAC2)) gene ESTs, STSs, GSSs and CpG islands,	Mycobacterium leprae Homo sapiens	61,170 37,455	27-Aug-99 16-Jun-99
rxa00799 1767	GB PL2:AF016327	616	AF016327	Hordeum vulgare Barperm1 (perm1) mRNA, partial cds.	Hordeum vulgare	41,311	01-OCT-1997
	GB_HTG2:HSDJ319M7 128208	7 128208		Homo sapiens chromosome 6 clone RP1-319M7 map p21.1-21.3, *** SEQUENCING IN PROGRESS *** in unordered pieces.	Homo sapiens	36,845	30-Nov-99
	GB_HTG2:HSDJ319M7 128208 AL079341	7 128208	AL079341	Homo sapiens chromosome 6 clone RP1-319M7 map p21.1-21.3, *** SEQUENCING IN PROGRESS *** in unordered pieces.	Homo sapiens	36,845	30-Nov-99
rxa00800 1227	GB_BA1:MTV022	13025	AL021925	Mycobacterium tuberculosis H37Rv complete genome; segment 100/162.	Mycobacterium	63,101	17-Jun-98
	GB BA1:AB019513	4417	AB019513	Streptomyces coelicolor genes for alcohol dehydrogenase and ABC	tuberculosis Streptomyces coelicolor	41.312	13-Nov-98
				transporter, complete cds.		!	
	GB_PL1:SCSFAARP	2008	X68020	S.cerevisiae SFA and ARP genes.	Saccharomyces cerevisiae 36,288	9 36,288	29-Nov-94
rxa00825 1056	GB_BA1:MTY15C10	33050	Z95436	Mycobacterium tuberculosis H37Rv complete genome; segment 154/162.	Mycobacterium tuberculosis	39,980	17-Jun-98
	GB_BA1:MLCB2548	38916	AL023093	Mycobacterium leprae cosmid B2548.	Mycobacterium leprae	39,435	27-Aug-99
	GB_BA2:AF169031	1141	AF169031	Xanthomonas oryzae pv. oryzae putative sugar nucleotide	Xanthomonas oryzae pv.	46,232	14-Sep-99
rxa00871				epimerase/dehydratase gene, partial cds.	oryzae		
rxa00872 1077	GB_IN1:CEF23H12	35564	Z74472	Caenorhabditis elegans cosmid F23H12, complete sequence.	Caenorhabditis elegans	34,502	08-OCT-1999
	GB_HTG2:AC007263	167390	AC007263	Homo sapiens chromosome 14 clone BAC 79J20 map 14q31, *** SEQUENCING IN PROGRESS ***, 5 ordered pieces.	Homo sapiens	35,714	24-MAY-1999
	GB_HTG2:AC007263	167390	167390 AC007263	Homo sapiens chromosome 14 clone BAC 79J20 map 14q31, *** SEQUENCING IN PROGRESS ***, 5 ordered pieces.	Homo sapiens	35,714	24-MAY-1999
rxa00879 2241	GB_BA1:MTV049	40360	AL022021	Mycobacterium tuberculosis H37Rv complete genome; segment 81/162.	Mycobacterium tuberculosis	36,981	19-Jun-98
	GB_PL2:CDU236897	1827	AJ236897	Candida dubliniensis ACT1 gene, exons 1-2.	Candida dubliniensis	38,716	1-Sep-99
	GB_PL1:CAACT1A	3206	X16377	Candida albicans act1 gene for actin.	Candida albicans	36,610	10-Apr-93
xa00909 955	GB_BA2:AF010496	189370	AF010496	Rhodobacter capsulatus strain SB1003, partial genome.	Rhodobacter capsulatus	51,586	12-MAY-1998
	GB_BA1:RMPHA	7888	X93358	Rhizobium meliloti pha[A,B,C,D,E,F,G] genes.	Sinorhizobium meliloti	48,367	12-MAR-1999
	GB_EST16:C23528	317	C23528	C23528 Japanese flounder spleen Paralichthys olivaceus cDNA clone HB5(2), mRNA sequence.	Paralichthys olivaceus	41,640	28-Sep-99
rxa00913 2118	GB_HTG2:AC007734	188267	188267 AC007734	Homo sapiens chromosome 18 done hRPK 44_O_1 map 18, *** SEQUENCING IN PROGRESS ***, 18 unordered pieces.	Homo sapiens	34,457	5-Jun-99

	GB_HTG2:AC007734		188267 AC007734	Table 4 (continued) Homo sapiens chromosome 18 clone hRPK.44 O 1 map 18 ****	Homo sapiens	34,457	6-Unr-9
	GE ECT18.8478	907	0.000470				C
	GG_EST 16.767 US476		AA/ 034/ 0	VV-4405.1 Durangene mouse near (#357310) Mus musculus CDNA clone IMAGE:1224272 5', mRNA sequence.	MIUS MUSCUIUS	42,065	. 24-DEC-1997
rxa00945 1095	GB_HTG4:AC010351	220710	220710 AC010351	Homo sapiens chromosome 5 clone CITB-H1_2022B6, *** SEQUENCING IN PROGRESS ***, 68 unordered pieces.	Homo sapiens	36,448	31-OCT-1999
	GB_HTG4:AC010351	220710	220710 AC010351	Homo sapiens chromosome 5 clone CITB-H1_2022B6, *** SEQUENCING IN PROGRESS ***, 68 unordered pieces.	Homo sapiens	36,448	31-OCT-1999
	GB_BA1:MTCY05A6	38631	296072	Mycobacterium tuberculosis H37Rv complete genome; segment 120/162.	Mycobacterium tuberculosis	36,218	17-Jun-98
rxa00965							
rxa00999 1575	GB_PAT:E13660	1916	E13660	gDNA encoding 6-phosphogluconate dehydrogenase.	Corynebacterium olutamicum	98,349	24-Jun-98
	GB_BA1:MTCY359	36021	Z83859	Mycobacterium tuberculosis H37Rv complete genome; segment 84/162.	Mycobacterium	38,520	17-Jun-98
	GB_BA1:MLCB1788	39228	AL008609		Mycobacterium leprae	64,355	27-Aug-99
rxa01015 442	GB_BA1:MTV008	63033	AL021246	Mycobacterium tuberculosis H37Rv complete genome; segment 108/162.	Mycobacterium tuberculosis	39,860	17-Jun-98
	GB_BA1:MTV008	63033	AL021246	Mycobacterium tuberculosis H37Rv complete genome; segment 108/162.	Mycobacterium tuberculosis	39,120	17-Jun-98
rxa01025 1119	GB_BA1:SC7A1	32039	AL034447	Streptomyces coelicolor cosmid 7A1.	Streptomyces coelicolor	55,287	15-DEC-1998
	GB_BA1:MSGB1723CS 38477	S 38477	L78825	Mycobacterium leprae cosmid B1723 DNA sequence.	Mycobacterium leprae	56,847	15-Jun-96
	GB_BA1:MLCB637	44882	<b>Z</b> 99263	Mycobacterium leprae cosmid B637.	Mycobacterium leprae	56,676	17-Sep-97
rxa01048 1347	GB_BA2:AF017444	3067	AF017444	Sinorhizobium meliloti NADP-dependent malic enzyme (tme) gene, complete	Sinorhizobium meliloti	53,660	2-Nov-97
	GB_BA1:BSUB0013	218470	218470 Z99116	cds. Bacillus subtilis complete genome (section 13 of 21): from 2395261 to	Bacillus subtilis	37,255	26-Nov-97
				2613730.			
	GB_VI:HSV2HG52	154746	154746 Z86099		human herpesvirus 2	38,081	04-DEC-1998
1500 1500 1500	GB_R1GZ:AC002578	137855	137855 AC002518	nomo sapiens chromosome X cione bWXUZU, *** SEQUENCING IN PROGRESS ***, 11 unordered pieces.	Homo sapiens	35,647	2-Sep-97
	GB_HTG2:AC002518	131855	131855 AC002518	Homo sapiens chromosome X clone bWXD20, *** SEQUENCING IN	Homo sapiens	35,647	2-Sep-97
	GB_HTG2:AC002518		131855 AC002518		Homo sapiens	26,180	2-Sep-97
				PROGRESS ***, 11 unordered pieces.			
rxa01077 1494	GB_PR3:HSDJ653C5	85237	AL049743	Human DNA sequence from clone 653C5 on chromosome 1p21.3-22.3 Contains CA repeat(D1S435), STSs and GSSs, complete sequence.	Homo sapiens	36,462	23-Nov-99
	GB_BA1:ECU29579	72221	U29579	Escherichia coli K-12 genome; approximately 61 to 62 minutes.	Escherichia coli	41,808	1-Jul-95
	GB_BA1:ECU29579	72221	U29579	Escherichia coli K-12 genome; approximately 61 to 62 minutes.	Escherichia coli	36,130	1-Jul-95
xa01089 873	GB_GSS8:AQ044021	387	AQ044021	CIT-HSP-2318C18.TR CIT-HSP Homo sapiens genomic clone 2318C18, genomic survey sequence.	Homo sapiens	36,528	14-Jul-98

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14-Jul-98 14-Jul-98 07-DEC-1994 17-Jun-98	23-DEC-1996 19-Feb-98 12-MAR-1998	01-MAR-1997 10-Jul-98 01-DEC-1999 13-Aug-99	22-Aug-99 22-Aug-99 05-OCT-1998 27-OCT-1994	07-0C1-1997 (Rel. 52, Created) 09-MAR-1995 07-0CT-1997 (Rel. 52, Created) 27-0CT-1994
35,969 44,545 100,000 63,771	67,061 99,615 100,000	37,785 35,835 37,873 37,240	38,416 38,416 38,416 99,931	39,753 100,000 100,000 36,769
Homo sapiens Homo sapiens Corynebacterium glutamicum Mycobacterium tuberculosis		Fugu rubripes Homo sapiens Homo sapiens Homo sapiens	Homo sapiens Homo sapiens Drosophila melanogaster Corynebacterium	Corynebacterium glutamicum Mycobacterium leprae Corynebacterium glutamicum glutamicum glutamicum Hepatitis C virus
Table 4 (continued) CIT-HSP-2318D17.TR CIT-HSP Homo sapiens genomic clone 2318D17, genomic survey sequence. CIT-HSP-2318C18.TR CIT-HSP Homo sapiens genomic clone 2318C18, genomic survey sequence. Corynebacterium pyruvate kinase gene, complete cds. Mycobacterium tuberculosis H37Rv complete genome; segment 72/162.	Mycobacterium intracellulare pyruvate kinase (pykF) gene, complete cds. Corynebacterium glutamicum inositol monophosphate phosphatase (impA) gene, complete cds. Corynebacterium glutamicum phosphoribosylformimino-5-amino-1-phosphoribosyl-4- imidazolecarboxamide isomerase (hisA) gene, complete cds.	F.rubripes GSS sequence, clone 079B16aE8, genomic survey sequence.  Homo sapiens chromosome 4 clone B32l8, complete sequence.  Human DNA sequence from clone RP5-1178l21 on chromosome X, complete sequence.  Homo sequence.  Homo sapiens clone NH0062F14, *** SEQUENCING IN PROGRESS ***, 5	Homo sapiens clone 1_O_3, *** SEQUENCING IN PROGRESS ***, 8 unordered pieces. Homo sapiens clone 1_O_3, *** SEQUENCING IN PROGRESS ***, 8 unordered pieces. Drosophila melanogaster cosmid 66A1. C.glutamicum (ASO 19) ATPase beta-subunit gene.	Mycobacterium leprae cosmid L471.  Mycobacterium leprae cosmid L471.  Brevibacterium flavum UncD gene whose gene product is involved in  C.glutamicum (ASO 19) ATPase beta-subunit gene.  Hepatitis C genomic RNA for putative envelope protein (RE4B isolate).
AQ042907 AQ044021 L27126 Z95554	U65430 AF045998 AF051846	619 Z89313 177014 AC004063 62268 AL109852 163369 AC009301	164587 AC009444 164587 AC009444 34127 AL031227 1452 X76875	U15186 U15186 E09634 X76875
392 387 2795 35938	1439 780 738	619 177014 62268 163369	164587 164587 34127 1452	1452 1452 1452 1452 414
GB_GSS8:AQ042907 GB_GSS8:AQ044021 GB_BA1:CORPYKI GB_BA1:MTCY01B2		GB_GSS1:FR0005503 GB_PR3:AC004063 GB_PR3:HS1178l21 GB_HTG3:AC009301	GB_HTG3:AC009444 GB_HTG3:AC009444 GB_IN1:DMC66A1 GB_BA1:CGASO19	GB_BA1:MLU15186 EM_PAT:E09634 GB_BA1:CGASO19 GB_VI:HEPCRE4B
rxa01093 1554	rxa01099 948	1111 541 xa01111	rxa01130 687	rxa01194 495

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01-MAY-1995	17-Jun-98	09-MAR-1995	01-MAY-1995	01-MAY-1995	26-MAY-1998	15-Sep-99	04-DEC-1999	04-DEC-1999	17-Jun-98	10-MAR-1998	26-Apr-93	2-Jun-98	17-Aug-99	17-Aug-99	13-MAR-1996	17-Jun-98	10-DEC-1996	09-OCT-1998 25-Sep-99	25-Sep-99
69,269	65,437	39,302	57,087	38,298	37,626	38,395	35,459	36,117	39,064	42,671	41,054	36,205	39,922	39,922	64,908	62,838	61,712	35,373 39,863	39,863
Streptomyces lividans	Mycobacterium tuberculosis	Mycobacterium leprae	Streptomyces lividans	Streptomyces lividans	Methylococcus capsulatus 37,626	Chloroplast Arabidopsis thaliana	Homo sapiens	Homo sapiens	Mycobacterium tuberculosis	Methylobacterium extorquens	Caulobacter crescentus	Streptomyces roseofulvus	Drosophila melanogaster	Drosophila melanogaster	Saccharopolyspora erythraea	Mycobacterium tuberculosis	Mycobacterium tuberculosis	Homo sapiens Homo sapiens	Homo sapiens
Table 4 (continued) S.lividans i protein and ATP synthase genes.	Mycobacterium tuberculosis H37Rv complete genome; segment 57/162.	Mycobacterium leprae cosmid L471.	S.lividans i protein and ATP synthase genes.	S.lividans i protein and ATP synthase genes.	M.capsulatus orfx, orfy, orfz, sqs and shc genes.	Arabidopsis thaliana chloroplast genomic DNA, complete sequence, strain:Columbia.	Homo sapiens clone RP11-114116, *** SEQUENCING IN PROGRESS ***, 39 unordered pieces.	Homo sapiens clone RP11-114I16, *** SEQUENCING IN PROGRESS ***, 39 unordered pieces.	Mycobacterium tuberculosis H37Rv complete genome; segment 47/162.	Methylobacterium extorquens methanol oxidation genes, glmU-like gene, partial cds, and orfL2, orfL1, orfR genes, complete cds.	C.crescentus flagellar gene promoter region.	Streptomyces roseofulvus frenolicin biosynthetic gene cluster, complete sequence.	Drosophila melanogaster chromosome 2 clone BACR04B09 (D576) RPCI-98 04.B.9 map 43E12-44F1 strain y; cn bw sp, *** SEQUENCING IN PROGRESS ***, 150 unordered pieces.	Drosophila melanogaster chromosome 2 clone BACR04B09 (D576) RPCI-98 04.B.9 map 43E12-44F1 strain y; cn bw sp, *** SEQUENCING IN PROGRESS ***, 150 unordered pieces.	Saccharopolyspora erythraea ferredoxin (fdxA) gene, complete cds.	Mycobacterium tuberculosis H37Rv complete genome; segment 51/162.	Mycobacterium tuberculosis sequence from clone y348.	Homo sapiens chromosome 17, clone hRPK.138_P_22, complete sequence. Homo sapiens clone NH0122L09, *** SEQUENCING IN PROGRESS ***, 2	unordered pieces. Homo sapiens clone NH0122L09, *** SEQUENCING IN PROGRESS ***, 2 unordered pieces.
Z22606	Z73419	U15186	Z22606	Z22606	Y09978	AP000423	164070 AC009762	164070 AC009762	<b>Z</b> 92539	AF017435	M69228	AF058302	165741 AC007301	165741 AC007301	M61119	AL010186	AD000020	AC005697 AC010722	160723 AC010722
8560	35516	36241	8560	8560	5538	154478	164070	164070	38970	4301	4424	25306	165741	165741	3869	37840	40056	174503 160723	160723
GB_BA1:SLATPSYNA	GB_BA1:MTCY373	GB_BA1:MLU15186	GB_BA1:SLATPSYNA	GB_BA1:SLATPSYNA	GB_BA1:MCSQSSHC	GB_PL1:AP000423	GB_HTG6:AC009762	GB_HTG6:AC009762	GB_BA1:MTCY10G2	GB_BA2:AF017435	GB_BA1:CCRFLBDBA 4424	GB_BA2:AF058302	GB_HTG3:AC007301	GB_HTG3:AC007301	GB_BA1:SERFDXA	GB_BA1:MTV005	GB_BA1:MSGY348	GB_PR3:AC005697 GB_HTG3:AC010722	GB_HTG3:AC010722
rxa01201 1764			rxa01202 1098			rxa01204 933			xa01216 1124			rxa01225 1563			rxa01227 444			rxa01242 900	

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23-OCT-1998	23-Nov-98	1-Feb-99	16-OCT-1999	16-OCT-1999	29-OCT-1999	5-Nov-97	21-MAR-1997	18-Apr-98	25-Sep-99	25-Sep-99	12-Nov-98 23-Jun-99	01-MAR-1994	2-Nov-99	2-Nov-99	2-Nov-99	07-DEC-1999	07-DEC-1999
38,722	35,448	100,000	37,178	37,178	59,719	59,735	37,904	37,340	36,385	36,385	39,494 46,252	46,368	36,016	36,016	39,618	35,366	35,366
Magnaporthe grisea	Caenorhabditis elegans	Corynebacterium	glutamicum Drosophila melanogaster	Drosophila melanogaster	Escherichia coli	Escherichia coli	Escherichia coli	Homo sapiens	Homo sapiens	Homo sapiens	Escherichia coli Mycobacterium tuberculosis	Mycobacterium leprae	Homo sapiens	Homo sapiens	Homo sapiens	Orosophila melanogaster	Drosophila melanogaster
mgxb0008N01r CUGi Rice Blast BAC Library Magnaporthe grisea genomic	cone nigatocono n. genomic suivey sequence. Caenorhabditis eligans coxxim (KOSD4, complete sequence.	Corynebacterium glutamicum lpd gene, complete CDS.	Drosophila melanogaster chromosome 3L/69C1 clone RPC198-11N6, ***	SECUENCING IN PROGRESS ***, /0 Unordered pieces.  Drosophila melanogaster chromosome 3L/69C1 clone RPCi98-11N6, ***SEQUENCING IN PROGRESS ***, 70 unordered pieces.	Escherichia coli GalF (galF) gene, partial cds; O-antigen repeat unit transporter Escherichia coli Wzx (wzx), WbnA (wbnA), O-antigen polymerase Wzy (wzy), WbnB (wbnB), WbnC (wbnC), WbnD (wbnD), WbnE (wbnE), UDP-Gic-4-epimerase GalE (galE), 6-phosphogluconate dehydrogenase Gnd (gnd), UDP-Gic-6-dehydrogenase Ugd (ugd), and WbnF (wbnF) genes, complete cds; and chain length dehydrogenase Ugd (wzy (wzy), nane nartial cds.	Escherichia communication (1975) going parameter and communication (1976) Escherichia coli Escherichia coli endo hypothetical uridina-65-diphosphoglucose dehydrogenase (1976) Escherichia coli and O-chain length regulator (1977) anne committee cas	E.coli genomic DNA, Kohara clone #351(45.1-45.5 min.).	Homo sapiens Xp22 BAC GS-619J3 (Genome Systems Human BAC library) complete sequence.	Homosapiens clone NH0310K15, *** SEQUENCING IN PROGRESS ***, 4	Homosapiens clone NH0310K15, *** SEQUENCING IN PROGRESS ***, 4 unordered pieces.	Escherichia coli K-12 MG1655 section 377 of 400 of the complete genome. Mycobacterium tuberculosis H37Rv complete genome; segment 143/162.	Mycobacterium leprae cosmid L308.	Homo sapiens chromosome 7, *** SEQUENCING IN PROGRESS ***, 24	Homosapiens chromosome 7, *** SEQUENCING IN PROGRESS ***, 24 unordered pieces.	Homo sapiens chromosome 7, *** SEQUENCING IN PROGRESS ***, 24 unordered pieces.	Drosophila melanogaster chromosome 2 clone BACR03D06 (D569) RPCI-98 03.D.6 map 32A-32A strain y. cn bw sp. *** SEQUENCING IN PROGRESS*** 91 unordered pieces.	Drosophila melanogaster chromosome 2 clone BACR19N18 (D572) RPCI-98 19.N.18 map 32A-32A strain y; cn bw sp, *** SEQUENCING IN PROGRESS ***, 22 unordered pieces.
AQ255057	Z92804 Z92804	Y16642	AC010567	143287 AC010567	AF172324	U78086	D90841	144368 AC004103	215529 AC007383	AC007383	AE000487 AL021841	U00022	215767 AC009245	215767 AC009245	AC009245	225851 AC007186	202291 AC007147
583	19000	1800	143287	143287	14263	4759	20226	144368	215529	215529	13889 53662	36411	215767	215767	215767	225851	202291
GB_GSS10:AQ255057	GB_IN1:CEK05D4	GB_BA1:CGLPD	GB_HTG4:AC010567	GB_HTG4:AC010567	GB_BA2:AF172324	GB_BA2:ECU78086	GB_BA1:D90841	GB_PR3:AC004103	GB_HTG3:AC007383	GB_HTG3:AC007383	GB_BA2:AE000487 GB_BA1:MTV016	GB_BA1:U00022	GB_HTG4:AC009245	GB_HTG4:AC009245	GB_HTG4:AC009245	GB_HTG6.AC007186	GB_HTG6:AC007147
rxa01243 1083		rxa01259 981			ка01262 1284			rxa01311 870			xa01312 2142		rxa01325 795			rxa01332 576	

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NHIBITORY FACTOR (HUMAN);, mRNA sequence.

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30-Jul-96	7-Aug-98	7-Aug-98	7-Aug-98	7-Aug-98	29-Jan-99	18-Sep-98	5-Aug-98	14-Jul-99	22-Jul-99	14-Jul-99 19-OCT-1999	19-OCT-1999	26-Jun-98	26-Nov-97	26-Nov-97	2-Aug-99
58,384	57,500	35,655	57,843	38,119	37,115	34,559	40,351	34,298	34,298	34,298 33,812	33,812	36,111	36,591	34,941	37,037
Neisseria meningitidis	Streptomyces griseus	Streptomyces griseus	Streptomyces griseus	Streptomyces griseus	Choristoneura fumiferana nucleopolyhedrovirus	Homo sapiens	Homo sapiens	Homo sapiens	Homo sapiens	Homo sapiens Drosophila melanogaster	Drosophila melanogaster	Homo sapiens	Bacillus subtilis	Bacillus subtilis	Drosophila melanogaster
Neisseria meningitidis dTDP-D-glucose 4,6-dehydratase (rbB), glucose-1-phosphate thymidyl transferase (rbA) and rbC genes, complete cds and UPD-phosphate thymidyl complete cds and UPD-phosphate	glucose-4-epimerase (galic) pseudogene. Streptomyces griseus genes for Orf2, Orf3, Orf4, Orf5, AfsA, Orf8, partial and complete cds.	Streptomyces griseus genes for Orf2, Orf3, Orf4, Orf5, AfsA, Orf8, partial and complete cds.	Streptomyces griseus genes for Orf2, Orf3, Orf4, Orf5, AfsA, Orf8, partial and complete cds.	Streptomyces griseus genes for Orf2, Orf3, Orf4, Orf5, AfsA, Orf8, partial and complete cds.	Choristoneura fumiferana nuclear polyhedrosis virus ETM protein homolog, 79 kDa protein homolog, 15 kDa protein homolog and GTA protein homolog qenes, complete cds.	HS_3249_B1_A02_MR CIT Approved Human Genomic Sperm Library D Homo Homo sapiens sapiens genomic clone Plate=3249 Col=3 Row=B, genomic survey sequence.	HS_3027_B1_H02_MR CIT Approved Human Genomic Sperm Library D Homo Homo sapiens sapiens genomic clone Plate=3027 Col=3 Row=P, genomic survey sequence.	Homo sapiens protocadherin gamma A3 short form protein (PCDH-gamma-A3) Homo sapiens variable region sequence, complete cds.	Homo sapiens protocadherin gamma A3 (PCDH-gamma-A3) mRNA, complete ods.	Homo sapiens PCDH-gamma-A3 gene, aberrantly spliced, mRNA sequence. Drosophila melanogaster chromosome 2 done BACR13N02 (D543) RPCI-98 13.N.2 map 36E-36E strain y; cn bw sp, *** SEQUENCING IN PROGRESS***, 101 unordered pieces.	Drosophila melanogaster chromosome 2 clone BACR13N02 (D543) RPCI-98 13.N.2 map 36E-36E strain y; cn bw sp, *** SEQUENCING IN PROGRESS***, 101 unordered bisces.	CIT-HSP-2280113.TR CIT-HSP Homo sapiens genomic clone 2280113, denomic survey sequence	Bacillus subtilis complete genome (section 9 of 21): from 1598421 to 1807200.	Bacillus subtilis complete genome (section 9 of 21): from 1598421 to 1807200.	Drosophila melanogaster chromosome 2 clone BACR48110 (D505) RPCI-98 48.I.10 map 49E6-49F8 strain y; cn bw sp, *** SEQUENCING IN PROGRESS ***, 17 unordered pieces.
L09189	AB011413	AB011413	AB011413	AB011413	U72240	AQ213248	AQ070145	AF152510	AF152323	AF152509 AC006590	127171 AC006590	B99182	299112	299112	174368 AC006247
8905	12070	12070	12070	12070	4783	408	285	2490	4605	2712 127171	127171	415	208780	208780	174368
GB_BA2:NGOCPSPS	GB_BA1:AB011413	GB_BA1:AB011413	GB_BA1:AB011413	GB_BA1:AB011413	GB_VI:CFU72240	GB_GSS10:AQ213248 408	GB_GSS8:AQ070145	GB_PR4:AF152510	GB_PR4:AF152323	GB_PR4:AF152509 GB_HTG4:AC006590	GB_HTG4:AC006590	GB_GSS8:B99182	GB_BA1:BSUB0009	GB_BA1:BSUB0009	GB_HTG2:AC006247
	rxa01571 723		rxa01572 615		rxa01606 2799			rxa01626 468		xa01632 1128			rxa01633 1206		

				Table 4 (continued)			
rxa01695 1623	GB_BA1:CGA224946	2408	AJ224946	Corynebacterium glutamicum DNA for L-Malate:quinone oxidoreductase.	Corynebacterium glutamicum	100,000	11-Aug-98
	GB_BA1:MTCY24A1	20270	295207	Mycobacterium tuberculosis H37Rv complete genome; segment 124/162.	Mycobacterium tuberculosis	38,626	17-Jun-98
rxa01702 1155	GB_IN1:DMU15974 GB_BA1:CGFDA	2994 3371	U15974 X17313	Drosophila melanogaster kinesin-like protein (klp68d) mRNA, complete cds. Corynebacterium glutamicum fda gene for fructose-bisphosphate aldolase (EC (	Drosophila melanogaster Corynebacterium	36,783 99,913	18-Jul-95 12-Sep-93
	GB_BA1:MTY13E10	35019	Z95324	terium tuberculosis H37Rv complete genome; segment 18/162.	giutamicumi Mycobacterium tuberculosis	38,786	17-Jun-98
	GB_BA1:MLCB4	36310	AL023514	Mycobacterium leprae cosmid B4.	Mycobacterium leprae	38,238	27-Aug-99
xa01743 901	GB_IN2:CELC27H5	35840	U14635		Caenorhabditis elegans	35,334	13-Jul-95
٠	GB_EST24:AI167112	629	Al167112	xylem.est.878 Poplar xylem Lambda ZAPII library Populus balsamifera subsp. F frichocama cDNA 5, mRNA sequence	Populus balsamifera subsolitrichocama	39,222	03-DEC-1998
	GB_GSS9:AQ102635	347	AQ102635	Human Genomic Sperm Library D Homo II=15 Row=L, genomic survey sequence.	Homo sapiens	40,653	27-Aug-98
rxa01744 1662	GB_BA1:MTCY01B2	35938	295554	Mycobacterium tuberculosis H37Rv complete genome; segment 72/162.	Mycobacterium tuberculosis	36,650	17-Jun-98
	GB_GSS1:AF009226	965	AF009226	Mycobacterium tuberculosis cytochrome D oxidase subunit I (appC) gene, partial sequence, genomic survey sequence.	Mycobacterium tuberculosis	63,438	31-Jul-97
	GB_BA1:SCD78	36224	AL034355		Streptomyces coelicolor	53,088	26-Nov-98
rxa01745 836	GB_BA1:MTCY190	34150	Z70283	complete genome; segment 98/162.	Mycobacterium tuberculosis	62,081	17-Jun-98
	GB_BA1:MLCB22	40281	298741	Mycobacterium leprae cosmid B22.	Mycobacterium leprae	61,364	22-Aug-97
	GB_BA2:AE000175	15067	AE000175	tion 65 of 400 of the complete genome.	Escherichia coli	52,323	12-Nov-98
rxa01758 1140	GB_PR3:HS57G9	113872	295116	Human DNA sequence from BAC 57G9 on chromosome 22q12.1 Contains 1 ESTs, CA repeat, GSS.	Homo sapiens	39,209	23-Nov-99
	GB_PL2:YSCH9666	39057	U10397	Saccharomyces cerevisiae chromosome VIII cosmid 9666.	Saccharomyces cerevisiae 40,021	40,021	5-Sep-97
	GB_PL2:YSCH9986	41664	U00027	Saccharomyces cerevisiae chromosome VIII cosmid 9986.	Saccharomyces cerevisiae 34,375	34,375	29-Aug-97
rxa01814 1785	GB_BA1:ABCCELB	2058	L24077	Acetobacter xylinum phosphoglucomutase (celB) gene, complete cds.	Acetobacter xylinus	62,173	21-Sep-94
	GB_BA1:MTCY22D7	31859	Z83866	Mycobacterium tuberculosis H37Rv complete genome; segment 133/162.	Mycobacterium tuberculosis	39,749	17-Jun-98
	GB_BA1:MTCY22D7	31859	Z83866	Mycobacterium tuberculosis H37Rv complete genome; segment 133/162.	Mycobacterium tuberculosis	40,034	17-Jun-98
تم1851 1809	GB_GSS9:AQ142579	529	AQ142579	HS_2222_B1_H03_MR CIT Approved Human Genomic Sperm Library D Homo Homo sapiens sapiens genomic clone Plate=2222 Col=5 Row=P, genomic survey sequence.	Homo sapiens	38,068	24-Sep-98
	GB_IN2:AC005889	108924	108924 AC005889	Drosophila melanogaster, chromosome 2L, region 30A3-30A6, P1 clones DS06958 and DS03097, complete sequence.	Drosophila melanogaster	36,557	30-OCT-1998
	GB_GSS1:AG008814	637	AG008814	Homo sapiens genomic DNA, 21q region, clone: B137B7BB68, genomic survey Homo sapiens sequence.	Homo sapiens	35,316	7-Feb-99

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	03-OCT-1999	15-Nov-99	03-OCT-1999	13-MAR-1996	17-Jun-98	10-DEC-1996	27-Apr-93 23-Nov-98	197	17-Jun-98	29-MAR-1999	7-Feb-99	20-Apr-99	2-Jun-99 28-Nov-98	03-DEC-1999	03-DEC-1999	01-OCT-1999	2-Aug-99
	36,364	35,334	36,529	59,862	61,949	59,908	36,899 36,899	34,805	37,892	40,413	47,792	43,231	39,306 42,807	36,417	37,667	39,640	32,969
	Microcystis aeruginosa	Trypanosoma brucei	Microcystis aeruginosa	Saccharopolyspora erythraea	Mycobacterium tuberculosis	Mycobacterium tuberculosis	Homo sapiens	Homo sapiens	Mycobacterium tuberculosis	Streptomyces coelicolor	Synechocystis sp.	nomo sapiens	Thermotoga maritima Drosophila melanogaster	Homo sapiens	Homo sapiens	Acinetobacter Iwoffii	Drosophila melanogaster
Table 4 (continued)	Microcystis aeruginosa DNA polymerase III beta subunit (dnaN) gene, partial cds; microcystin synthetase gene cluster, complete sequence; Uma1 (uma1), Uma2 (uma2), Uma3 (uma3), Uma4 (uma4), and Uma5 (uma5) genes, complete cds; and Uma6 (uma6) gene, partial cds.	Trypanosoma brucei chromosome II clone RPCI93-25N14, *** SEQUENCING IN PROGRESS ***, 2 unordered pieces.	Microcystis aeruginosa DNA polymerase III beta subunit (dnaN) gene, partial cds; microcystin synthetase gene cluster, complete sequence; Uma1 (uma1), Uma2 (uma2), Uma3 (uma3), Uma4 (uma4), and Uma5 (uma5) genes, complete cds; and Uma6 (uma6) gene, partial cds.	Saccharopolyspora erythraea ferredoxin (fdxA) gene, complete cds.	Mycobacterium tuberculosis H37Rv complete genome; segment 51/162.	Mycobacterium tuberculosis sequence from clone y348.	Human kidney alpha-2-adrenergic receptor mRNA, complete cds. Homo saniens alpha2-C4-adreneraic receptor cane complete cds.	HS-1055-B1-A03-MR abi CIT Human Genomic Sperm Library C Homo sapiens Homo sapiens genomic clone Plate=CT 777 Col=5 Row=B, genomic survey sequence.	Mycobacterium tuberculosis H37Rv complete genome; segment 69/162.	Streptomyces coelicolor A3(2), glycogen metabolism cluster II.	Synechocystis sp. PCC6803 complete genome, 10/27, 1188886-1311234.	KTCI I 44TO. IK. I KTCI I I nomo sapiens genomic cione KTCI I 44TO, genomic survey sequence.	Thermotoga maritima section 33 of 136 of the complete genome. GM01044.5prime GM Drosophila melanogaster ovary BlueScript Drosophila melanogaster cDNA clone GM01044 5prime. mRNA sequence	Homo sapiens clone RP3-405J10, *** SEQUENCING IN PROGRESS ***, 102	unordered pleces.  Homo sapiens clone RP3-405J10, *** SEQUENCING IN PROGRESS ***, 102 unordered pieces.	Activation process. Activate with week, week, week, week, wzy, week, week, week, week, week, week, week, week, week, galu, ugd, pgi, galk, pgm (partial) and mip (partial) genes (emulsan biosynthetic gene cluster), strain PAG-1.	Drosophila melanogaster chromosome 3 clone BACR02L12 (D753) RPCI-98 02.L.12 map 94B-94C strain y; cn bw sp, *** SEQUENCING IN PROGRESS*** 113 unordered pieces.
	AF183408	158889 AC008031	AF183408	M61119	AL010186	AD000020	J03853 U72648	B42200	274020	AJ001206		AQ116291	AE001721 AA567090	303147 AC008147	303147 AC008147	AJ243431	125235 AC008197
	63626	158889	63626	3869	37840	40056	1491	387	35377	9184	122349	7/6	17632 596	303147	303147	26953	125235
	GB_BA2:AF183408	GB_HTG5:AC008031	GB_BA2:AF183408	GB_BA1:SERFDXA	GB_BA1:MTV005	GB_BA1:MSGY348	GB_PR1:HUMADRA2C 1491	GB_GSS3:B42200	GB_BA1:MTCY48	GB_BA1:SCO001206	GB_BA1:D90908	167911DY:6005	GB_BA2:AE001721 GB_EST16:AA567090	GB_HTG6:AC008147	GB_HTG6:AC008147	GB_BA2:ALW243431	GB_HTG2:AC008197
	rxa01859 1050			rxa01865 438			xa01882 1113		xa01884 1913		000000	1xa01060 69/		rxa01887 1134			rxa01888 658

				Table 4 (continued)			
	GB_HTG2:AC008197	125235	125235 AC008197	Drosophila melanogaster chromosome 3 clone BACR02L12 (D753) RPCI-98	Drosophila melanogaster	32,969	2-Aug-99
				***, 113 unordered pieces.			
	GB_EST36:AI881527	298	AI881527	606070C09.y1 606 - Ear tissue cDNA library from Schmidt lab Zea mays cDNA, Zea mays mRNA sequence.	, Zea mays	43,617	21-Jul-99
гха01891 887	GB_VI:HIV232971	621	AJ232971	Human immunodeficiency virus type 1 subtype C nef gene, patient MP83.	Human immunodeficiency	40,040	05-MAR-1999
	GB_PL1:AFCHSE	6158	Y09542	A.fumigatus chsE gene.	Aspergillus fumigatus	37,844	1-Apr-97
	GB_PR3:AF064858	193387	AF064858	Homo sapiens chromosome 21q22.3 BAC 28F9, complete sequence.	Homo sapiens	37,136	2-Jun-98
xa01895 1051	GB_BA1:CGL238250	1593	AJ238250	Corynebacterium glutamicum ndh gene.	Conynebacterium	100,000	24-Apr-99
	GB_BA2:AF038423	1376	AF038423	Mycobacterium smegmatis NADH dehydrogenase (ndh) gene, complete cds.	glutamicum Mycobacterium smegmatis 65,254	65,254	05-MAY-1998
	GB_BA1:MTCY359	36021	Z83859	Mycobacterium tuberculosis H37Rv complete genome, segment 84/162.	Mycobacterium	40,058	17-Jun-98
CAD1401 1383	GB BA1-MSGB38COS	37114	101005	M tense appoint DNA contense seemid R18 hfrage appointed and	(uberculosis Mycobacterium leome	50 551	S. Con 04
	GB_BA1:SCE63 37200	37200	AL035640	Streptomyces coelicolor cosmid E63.	Streptomyces coelicolor	39,468	17-MAR-1999
	GB_PR3:AF093117	147216	AF093117	Homo sapiens chromosome 7qtelo BAC E3, complete sequence.	Homo sapiens	39,291	02-OCT-1998
rxa01927 1503	GB_BA1:CGPAN	2164	X96580	C.glutamicum panB, panC & xylB genes.	Corynebacterium	38,384	11-MAY-1999
	GB_BA1:ASXYLA	1905	X59466	Arthrobacter Sp. N.R.R.L. B3728 xylA gene for D-xylose(D-glucose) isomerase. Arthrobacter sp.	Arthrobacter sp.	56,283	04-MAY-1992
	GB_HTG3.AC009500	176060	176060 AC009500	Homo sapiens clone NH0511A20, *** SEQUENCING IN PROGRESS ***, 6 unordered pieces.	Homo sapiens	37,593	24-Aug-99
rxa01952 1836	GB_BA2:AE000739	13335	AE000739	Aquifex aeolicus section 71 of 109 of the complete genome.	Aquifex aeolicus	36,309	25-MAR-1998
	GB_EST28:AI519629	612	AI519629	LD39282.5prime LD Drosophila melanogaster embryo pOT2 Drosophila melanogaster cDNA clone LD39282 5prime, mRNA sequence.	Drosophila melanogaster	41,941	16-MAR-1999
	GB_EST21:AA949396	792	AA949396	LD28277.5prime LD Drosophila melanogaster embryo pOT2 Drosophila melanogaster cDNA clone LD28277 5prime, mRNA sequence.	Drosophila melanogaster	39,855	25-Nov-98
rxa01989 630	GB_BA1:BSPGIA	1822	X16639	Bacillus stearothermophilus pgiA gene for phosphoglucoisomerase isoenzyme A (EC 5.3.1.9).	Bacillus stearothermophilus	66,292	20-Apr-95
	GB_BA1:BSUB0017	217420	217420 Z99120	Bacillus subtilis complete genome (section 17 of 21); from 3197001 to 3414420.	Bacillus subtilis	37,255	26-Nov-97
	GB_BA2:AF132127	8452	AF132127	Streptococcus mutans sorbitol phosphoenolpyruvate:sugar phosphotransferase Streptococcus mutans operon, complete sequence and unknown gene.	Streptococcus mutans	63,607	28-Sep-99
rxa02026 720	GB_BA1:SXSCRBA	3161	X67744	S.xylosus scrB and scrR genes.	Staphylococcus xylosus	67,778	28-Nov-96
	GB_BA1:BSUB0020	212150	Z99123	Bacillus subtilis complete genome (section 20 of 21): from 3798401 to	Bacillus subtilis	35,574	26-Nov-97
	GB_BA1:BSGENR	97015	X73124	B.subtilis genomic region (325 to 333).	Bacillus subtilis	51,826	2-Nov-93
ra02028 526	GB_BA1:MTCI237	27030	294752	Mycobacterium tuberculosis H37Rv complete genome; segment 46/162.	Mycobacterium tuberculosis	54,476	17-Jun-98

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	1-Aug-97	29-Apr-99		27-Aug-99	17-Jun-98	14-Aug-97	28.11.69	) ;	6-Feb-99		18-Jun-98	08-DEC-1999		19-DEC-1997		04-MAR-1998	12-Jul-99	1-Jul-99		27-OCT-1999	10.DEC.1998	2000	17-Jun-98	20 848 0 1000	58-INIAR-1888	29-MAY-1997	65-deS-9	15-Sep-99	15-Jun-96	15-Jun-96	01-OCT-1999	17-Jun-98
	le 36,100	ie 32,039		61,896	59,964	59,659	800 80	200	98,928		39,265	37,453		37,711		37,711	56,972	40,696		36,795	40 156	2	55,218	20 475	56,475	38,586	37,259	38,868	51,399	51,399	36,683	57,292
	Saccharomyces cerevisiae 36,100	', Saccharomyces cerevisia		Mycobacterium leprae	Mycobacterium tuberculosis	Mycobacterium	Copynehacterium	glutamicum	e Corynebacterium	glutamicum	Mycobacterium	Homo sapiens		e Arabidopsis thaliana		· Arabidopsis thaliana	Streptomyces coelicolor	Onyza sativa		Homo sapiens	Machacharium	Inherculosis	Mycobacterium	tuberculosis	streptornyces coencolor	Escherichia coli	, Homo sapiens	Homo sapiens	Mycobacterium leprae	Mycobacterium leprae	Mus musculus	Mycobacterium tuberculosis
Table 4 (continued)	Saccharomyces cerevisiae chromosome V cosmids 9537, 9581, 9495, 9867,	and tanipua done 3050. V26G9 mTn-3xHA/lacZ Insertion Library Saccharomyces cerevisiae genomic 5', Saccharomyces cerevisiae 32,039	genomic survey sequence.	Mycobacterium leprae cosmid B1222.	Mycobacterium tuberculosis H37Rv complete genome; segment 147/162.	Mycobacterium tuberculosis rfbA, rhamnose biosynthesis protein (rfbA), and	rmic genes, complete cas. Revibacterium lactofermentum nene for sloba-ketochularic acid	dehydrogenase.	Corynebacterium glutamicum DNA for 2-oxoglutarate dehydrogenase, complete Corynebacterium	cds.	Mycobacterium tuberculosis H37Rv complete genome; segment 54/162.			_	sequence.	Arabidopsis thaliana chromosome II BAC F25I18 genomic sequence, complete Arabidopsis thaliana sequence.	S.coelicolor DNA for glgC gene.		nbxb0074H11r, genomic survey sequence.				Mycobacterium tuberculosis H37Ry complete genome; segment 59/162.	to the state of th	Sireptornyces coelicolor A3(z) glycogen metabolism ciusten.	E.coli genomic DNA, Kohara clone #401(51.3-51.6 min.).	wq07d12.x1 NCI_CGAP_Kid12 Homo sapiens cDNA clone IMAGE:2470583 3'. Homo sapiens mRNA sequence.	Homo sapiens chromosome 5 clone CITB-H1_2074D8, *** SEQUENCING IN PROGRESS ***, 77 unordered pieces.	Mycobacterium leprae cosmid B1551 DNA sequence.	Mycobacterium leprae cosmid B1554 DNA sequence.	Mus musculus transcription factor TBLYM (Tblym) mRNA, complete cds.	Mycobacterium tuberculosis H37Ry complete genome; segment 98/162.
	018778	AQ501177		AL049491	Z95390	U43540	F14601		D84102		AL021006	211682 AC005883		AC003033		AC002334	X89733	AQ687350		AW028530	AD00018		Z73902	100400	A3001203	D90858	AI948595	220665 AC010387	L78813			Z70283
	66030	767		34714	43401	3453	4394	}	4394		22440	211682		84254		75050	1518	786		444	37036	3	32514	0000	6006	13548	469	220665	36548	36548	2482	34150
	GB_PL2:SCE9537	GB_GSS13:AQ501177		GB_BA1:MLCB1222	GB_BA1:MTY13E12	GB_BA1:MTU43540	GR DAT:#14601		GB_BA1:D84102		GB_BA1:MTV006	GB_HTG7:AC005883	` I	GB_PL2:ATAC003033		GB_PL2:ATAC002334	GB_BA1:SCGLGC	GB_GSS4:AQ687350		GB_EST38:AW028530 444	CB BA1-MSGV151		GB_BA1:MTCY130	984.600004305	GB_BAT:SCUUTZUS	GB_BA1:D90858	GB_EST37:AI948595	GB_HTG3:AC010387	GB BA1:MSGB1551CS	GB_BA1:MSGB1554C	GB_RO:AF093099 2482	GB_BA1:MTCY190
				rxa02054 1140			rs02056 2891					xa02061 1617					rxa02063 1350				845C 001C0cx	14802.100 2540				rxa02122 822			rxa02140 1200			rxa02142 774

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	24-MAR-1999	5-Aug-99	17-Jun-98	15-Jun-96	15-Jun-96	17-Jun-98	18-Feb-00		18-Feb-00	30-MAR-1999		11-DEC-1996		11-DEC-1996	23-Nov-99		06-DEC-1999	2-Jul-99		17-Feb-95	;	17-Jun-98	10-Feb-99	31-MAY-1995	4-hin-98		31-MAY-1995		3-Apr-99	26-MAR-1998
	35,058	47,403	57,317	38,159	38,159	55,530	39,659		39,659	39.798	<u>.</u>	36,436		36,436	36,872		43,175	39,715		100,000		64,331	62,491	38,791	40 044		37,312		99,173	40,219
	Streptomyces coelicolor	Pseudomonas putida	Mycobacterium tuberculosis	Mycobacterium leprae	Mycobacterium leprae	Mycobacterium tuberculosis	Homo sapiens		Homo sapiens	Homo sapiens		Homo sapiens		Homo sapiens	Homo sapiens	-	Rhizobium etli	Mus musculus		Conynebacterium	glutamicum	Mycobacterium tuberculosis	Mycobacterium leprae	Rattus norvegicus	Rhodobacter enbaeroides		Rattus norvegicus		Corynebacterium glutamicum	Mycobacterium avium
Table 4 (continued)	Streptomyces coelicolor cosmid 6G10.	Pseudomonas putida genes for cytochrome o ubiquinol oxidase A-E and 2 ORFs, complete cds.	Mycobacterium tuberculosis H37Rv complete genome; segment 98/162.	Mycobacterium leprae cosmid B1551 DNA sequence.	Mycobacterium leprae cosmid B1554 DNA sequence.	Mycobacterium tuberculosis H37Rv complete genome; segment 98/162.	Homo sapiens chromosome 19 clone CIT978SKB_60E11, *** SEQUENCING	IN PROGRESS ***, 246 unordered pieces.	Homo sapiens chromosome 19 done CIT978SKB_60E11, *** SEQUENCING IN PROCRESS *** 246 unordered pieces	ta07a01.x1 NCI CGAP CLL1 Homo sapiens cDNA clone IMAGE:2108040 3'.	mRNA sequence.	zo50e01.r1 Stratagene endothelial cell 937223 Homo sapiens cDNA clone	IMAGE:590328 5', mRNA sequence.	zo50e01.r1 Stratagene endothelial cell 937223 Homo sapiens cDNA clone IMAGE:590328 5' mRNA sequence.	Human DNA sequence from clone 277P6 on chromosome 1q25.3-31.2,	complete sequence.	Rhizobium etli mutant MB045 RosR-transcriptionally regulated sequence.	uk53g05.y1 Sugano mouse kidney mkia Mus musculus cDNA clone	IMAGE:1972760 5' similar to WP:K11H12.8 CE12160;, mRNA sequence.	C.glutamicum glt gene for citrate synthase and ORF.		Mycobacterium tuberculosis H3/Rv complete genome; segment 41/162.	Mycobacterium leprae cosmid B57.	Rattus norvegicus dipeptidyl aminopeptidase-related protein (dpp6) mRNA,	complete cds. 127DB037070107 Cosmid library of chromosome II Bhodobarter subservides	genomic clone 127PB037070197, genomic survey sequence.	Rattus norvegicus dipeptidyl aminopeptidase-related protein (dpp6) mRNA,	complete cds.	Corynebacterium glutamicum gene for aconitase, partial cds.	Mycobacterium avium strain GIR10 transcriptional regulator (mav81) gene, partial cds, aconitase (acn), invasin 1 (inv1), invasin 2 (inv2), transcriptional regulator (moxR), ketoacyl-reductase (fabG), enoyl-reductase (inhA) and ferrochelatase (mav272) genes, complete cds.
	AL049497	AB016787	Z70283	L78813	L78814	Z70283	AC011500		AC011500	AI492095		AA157467		AA157467	AL117347		AF116423	AI789323		X66112		273101	<b>Z99494</b>	M76426	A0012162	30.4	M76426		AB025424	AF002133
	36734	5550	34150	36548	36548	34150	300851		300851	485	}	376		376	61698		360	574		3013		37630	38029	2819	763	3	2819		2995	15437
	GB_BA1:SC6G10	GB_BA1:AB016787	GB_BA1:MTCY190	GB_BA1:MSGB1551CS 36548	GB_BA1:MSGB1554CS 36548	GB_BA1:MTCY190	GB_HTG3:AC011500_0 300851		GB_HTG3:AC011500_0 300851	GB EST28:Al492095	•	GB_EST10:AA157467		GB_EST10:AA157467	GB PR3:HSBK277P6		GB_BA2:EMB065R075	GB_EST34:AI789323		GB_BA1:CGGLTG		GB_BA1:MTCY31	GB_BA1:MLCB57	GB_RO:RATDAPRP	GB GSC8:40012162	70171070707070	GB_RO:RATDAPRP		GB_BA1:AB025424	GB_BA2:AF002133
			xa02143 1011			rxa02144 1347				rxa02147 1140					rxa02149 1092					xa02175 1416				rxa02196 816					rxa02209 1694	

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38,253	960'66	34,937	36.885		48,701		39,119	33,118	99,289	<u> </u>	36,951	64,196	;	98,873	61,273	61 772		299'66	,	000'001	100,000	100,000		000'001	99,827
Mycobacterium	tuberculosis Corynebacterium	glutamicum Mycobacterium	tuberculosis Mycobacterium avium		Rhodobacter capsulatus		Escherichia coli	Homo sapiens	Corvnebacterium	glutamicum	Streptomyces coelicolor	Mycobacterium	tuberculosis	Corynebacterium	Mycobacterium	tuberculosis Mycobacterium avium		Corynebacterium .	glutamicum	Corynebacterium alutamicum	Conynebacterium	glutamicum Corynebacterium	glutamicum	Corynebacterium	gotamodin Corynebacterium glutamicum
AL021184 Mycobacterium tuberculosis H37Rv complete genome; segment 64/162.	Corynebacterium glutamicum gene for aconitase, partial cds.	Mycobacterium tuberculosis H37Rv complete genome; segment 64/162.	Mycobacterium avium strain GIR10 transcriptional regulator (may81) gene	parial cds, aconitase (acn), invasin 1 (inv1), invasin 2 (inv2), transcriptional regulator (moxR), ketoacyl-reductase (fabG), enoyl-reductase (inhA) and ferror-helatase (may 72) nanes complete cds	Rhodobacter capsulatus Calvin cycle carbon dioxide fixation operon: fructose-1,5-/sedoheptulose-1,7-bisphosphate aldolase (cbbA) gene, partial cds, Form II	ribulose-1,5-bisphosphate carboxylase/oxygenase (cbbM) gene, complete cds, and Calvin cycle operon: pentose-5-phosphate-3-epimerase (cbbE), phosphoglycolate phosphatase (cbbZ), and cbbY genes, complete cds.	Escherichia coli minutes 9 to 11 genomic sequence.	Homo sapiens chromosome 18 clone hRPK.178_F_10 map 18, *** centencials in poocees *** 11 moodead siess	SECCENCING IN PROGRESS 1.11 unblocked pieces.  Colutamicum gap, gok and toi genes for glyceraldehyde-3-phosphate.	phosphoglycerate kinase and triosephosphate isomerase.	Streptomyces coelicolor cosmid C54.	Mycobacterium tuberculosis H37Rv complete genome; segment 63/162.		C.glutamicum gap, pgk and tpi genes for glyceraldehyde-3-phosphate, phosphoponycerate kinase and triosenhosphate isomerase	Mycobacterium tuberculosis H37Rv complete genome; segment 63/162.	Mucobarteniim aviiim alvoeraldebude. 3. ahosahate debudronenase homolon	(gapdh) gene, complete cds; and phosphoglycerate kinase gene, partial cds.	C.glutamicum gap, pgk and tpi genes for glyceraldehyde-3-phosphate,	phosphoglycerate kinase and triosephosphate isomerase.	C.glutamicum phosphoenolpyruvate carboxylase gene, complete cds.	C.glutamicum ppg gene for phosphoenol pyruvate carboxylase.	C. clutamicum phosphoenolovruvate carboxvlase gene, complete cds.	-	C.glutamicum ppg gene for phosphoenol pyruvate carboxylase.	Corynebacterium glutamicum phosphoenolpyruvate carboxylase gene (EC 4.1.31).
AL021184	AB025424	AL021184	AF002133		U23145		139818 U82664	158858 AC007922	X59403		AL035591	<b>Z</b> 95844	;	X59403	Z95844	1182740		X59403		M25819	A09073	M25819		A09073	X14234
32806	2995	32806	15437		2960		139818	158858	3804		30753	40790		3804	40790	2530	}	3804		488 5	4885	4885	,	4885	3292
GB_BA1:MTV007	_ GB_BA1:AB025424	GB_BA1:MTV007	GB BA2:AF002133		GB_BA2:RCU23145		GB_BA1:ECU82664	GB_HTG2:AC007922	GB BA1:CGGAPPGK		GB_BA1:SCC54	GB_BA1:MTCY493		GB_BA1:CGGAPPGK	GB_BA1:MTCY493	GB BA2-MA1182740		GB_BA1:CGGAPPGK		GB_BA1:CORPEPC	GB_PAT:A09073	GB BA1:CORPEPC	1	68_PA1:A09073	GB_BA1:CGPPC
	1 874 X802213 874				rxa02245 780				rxa02256 1125					xa02257 1338				rxa02258 900				rxa02259 2895			

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	03-DEC-1999	11-Sep-99	•	11-Sep-99		01-DEC-1999	0	66-106-01	3	21-Apr-9/	17-Jun-98	17-Jun-98		08-MAY-1998		24-DEC-1997	17. hip.98		08-MAY-1998		24-DEC-1997	00	17-Jun-98	08-MAY-1998		24-DEC-1997	29-OCT-1998	20-Sep-91	23-Sep-97		12-Nov-98	20-Sep-91	23-Nov-99
	36,039	35,331		35,331		39,747	9	29, 100	100	35,922	57,677	37,143		100,000		100,000	17 163		99,259		99,259	1	/١٤,١4	100,000		100,000	52,248	tis 58,460	- 57,154		38,164	tis 58,929	33,070
	Homo sapiens	Homo sapiens	•	Homo sapiens		Bacteroides fragilis		nomo sapiens		Homo sapiens	Mycobacterium tuberculosis	Mycobacterium	tuberculosis	Corynebacterium	glutamicum	Corynebacterium	Mycobacterium	tuberculosis	Corynebacterium	glutamicum	Conynebacterium	glutamicum	Mycobacterium tuberculosis	Corynebacterium	glutamicum	Corynebacterium glutamicum	Aspergillus terreus	Mycobacterium smegmatis 58,460	Antarctic bacterium DS2-	3R	Escherichia coli	Mycobacterium smegmatis 58,929	Homo sapiens
Table 4 (continued)	Human DNA sequence from clone RP1-94E24 on chromosome 20q12,	complete sequence. Homo sapiens clone NH0295A01, *** SEQUENCING IN PROGRESS ***, 4	unordered pieces.	Homo sapiens clone NH0295A01, *** SEQUENCING IN PROGRESS ***, 4	unordered pieces.	Bacteroides fragilis 638R polysaccharide B (PS B2) biosynthesis locus,	complete sequence; and unknown genes.	no_pount along Distant 1084 Oct-11 Duman Maie BAC Library nomic saprens	genomic cione riade 1001 Cole 12 NOW-E, genomic survey sequence.	EST95058 Activated 1-cells I Homo sapiens cunA 5 end, mkNA sequence.	Mycobacterium tuberculosis H3/Rv complete genome; segment 49/162.	Mycobacterium tuberculosis H37Rv complete genome; segment 49/162.		Corynebacterium glutamicum pyc gene.		Corynebacterium glutamicum pyruvate carboxylase (pyc) gene, complete cds.	Avcohacterium fulberculosis H37Rv complete genome: segment 134/162		Corynebacterium glutamicum pyc gene.		Corynebacterium glutamicum pyruvate carboxylase (pyc) gene, complete cds.		Mycobacterium (uperculosis H3/RV complete genome; segment 131/162.	Corynebacterium glutamicum pyc gene.		Corynebacterium glutamicum pyruvate carboxylase (pyc) gene, complete cds.	Aspergillus terreus pyruvate carboxylase (Pyc) mRNA, complete cds.	M.smegmatis gltA gene for citrate synthase.	Antarctic bacterium DS2-3R citrate synthase (cisy) gene, complete cds.		Escherichia coli K-12 MG1655 section 65 of 400 of the complete genome.	M.smegmatis gltA gene for citrate synthase.	GB_PR4:HUAC002299 171681 AC002299 Homo sapiens Chromosome 16 BAC clone CIT987-SKA-113A6 ~complete genomic sequence, complete sequence.
	243145 AL050317	159526 AC010091		159526 AC010091		AF125164	703777	70/44695		AA381925	782282	295585		Y09548		AF038548	783018		Y09548		AF038548	10001	283018	Y09548		AF038548	AF097728	X60513	U85944		AE000175	X60513	AC002299
	243145	159526		159526		26443	3	/70	6	505	22550	22550		3728		3637	43523		3728	8	3637	101	43523	3728		3637	3916	1776	1334	1	15067	1776	171681
	GB_PR3:HSDJ94E24	GB_HTG3:AC010091	ı	GB_HTG3:AC010091		GB_BA2:AF125164	2007 A CO 4 CO CO	GB_G995.AQ7.44695	100100	GB_ES114:AA381925	GB_BA1:M1CY22G8	GB_BA1:MTCY22G8		GB_BA1:CGPYC		GB_BA2:AF038548	GR BA1-MTCY349		GB_BA1:CGPYC		GB_BA2:AF038548	000000000000000000000000000000000000000	66_6A1:M1C1349	GB_BA1:CGPYC		GB_BA2:AF038548	GB_PL2:AF097728	GB_BA1:MSGLTA	GB_BA2:ABU85944		GB_BA2:AE000175	GB_BA1:MSGLTA	GB_PR4:HUAC002299
	rxa02288 969					xa02292 798					rxa02322 511			xa02326 939					rxa02327 1083					rxa02328 1719				xa02332 1266				rxa02333 1038	

				Table 4 (continued)			
	GB_HTG2:AC007889	127840	127840 AC007889	Drosophila melanogaster chromosome 3 clone BACR48E12 (D695) RPCI-98 48 E 12 map 87A-878 strain v. on bw so SFOLIFINGIN PROGRESS.**	Drosophila melanogaster	34,897	2-Aug-99
				86 unordered pieces.			
rxa02399 1467	GB_BA1:CGACEA	2427	X75504	C.glutamicum aceA gene and thiX genes (partial).	Corynebacterium glutamicum	100,000	9-Sep-94
	GB_BA1:CORACEA	1905	128760	Corynebacterium glutamicum isocitrate lyase (aceA) gene.	Corynebacterium glutamicum	100,000	10-Feb-95
	GB_PAT:113693	2135	113693		Unknown.	99,795	26-Sep-95
rxa02404 2340	GB_BA1:CGACEB	3024	X78491	C.glutamicum (ATCC 13032) aceB gene.	Corynebacterium	99,914	13-Jan-95
	GB_BA1:CORACEB	2725	L27123	Corynebacterium glutamicum malate synthase (aceB) gene, complete cds.	glutamicum Corynebacterium	99,786	8-Jun-95
	GB BA1:PFFC2	5588	Y11998	Pilluorescens FC2.1 FC2.2 FC2.3c FC2.4 and FC2.5c open reading frames.	glutamicum Pseudomonas fluorescens 63,539	63.539	11-Jul-97
						200	
ra02414 870	GB_PR4:AC007102	176258	176258 AC007102	Homo sapiens chromosome 4 clone C0162P16 map 4p16, complete sequence. Homo sapiens	Homo sapiens	35,069	2-Jun-99
	GB_HTG3:AC011214	183414	183414 AC011214	Homo sapiens clone 5_C_3, LOW-PASS SEQUENCE SAMPLING.	Homo sapiens	36,885	03-OCT-1999
	GB_HTG3:AC011214	183414		Homo sapiens done 5_C_3, LOW-PASS SEQUENCE SAMPLING.	Homo sapiens	36,885	03-OCT-1999
rxa02435 681	GB_BA2:AF101055	7457	AF101055	Clostridium acetobutylicum atp operon, complete sequence.	Clostridium acetobutylicum 39,605	39,605	03-MAR-1999
	GB_OM:RABPKA		J03247	Rabbit phosphorylase kinase (alpha subunit) mRNA, complete cds.	Oryctolagus cuniculus	36,061	27-Apr-93
	GB_OM:RABPLASISM	4458	M64656	Oryctolagus cuniculus phosphorylase kinase alpha subunit mRNA, complete cds.	Oryctolagus cuniculus	36,000	22-Jun-98
rxa02440 963	GB_EST14:AA417723	374	AA417723	zv01b12.s1 NCI_CGAP_GCB1 Homo sapiens cDNA clone IMAGE:746207 3' similar to contains Alu repetitive element; contains element L1 repetitive element; mRNA sequence.	Homo sapiens	38,770	16-OCT-1997
	GB_EST11:AA215428	303	AA215428	zr95a07.s1 NCL_CGAP_GCB1 Homo sapiens cDNA clone IMAGE:683412 3' similar to contains Alu repetitive element; mRNA sequence.	Homo sapiens	39,934	13-Aug-97
	GB_BA1:MTCY77	22255	<b>Z</b> 95389	Mycobacterium tuberculosis H37Rv complete genome; segment 146/162.	Mycobacterium tuberculosis	38,889	18-Jun-98
rxa02453 876	GB_EST14:AA426336	375	AA426336	zv53g02.s1 Soares_testis_NHT Homo sapiens cDNA clone IMAGE:757394 3', mRNA sequence.	Homo sapiens	38,043	16-OCT-1997
	GB_BA1:STMAACC8	1353	M55426	S.fradiae aminoglycoside acetyltransferase (aacC8) gene, complete cds.	Streptomyces fradiae	37,097	05-MAY-1993
	GB_PR3:AC004500	77538	AC004500	Homo sapiens chromosome 5, P1 clone 1076B9 (LBNL H14), complete sequence.	Homo sapiens	33,256	30-MAR-1998
rxa02474 897	GB_BA1:AB009078	2686	AB009078	Brevibacterium saccharolyticum gene for L-2.3-butanediol dehydrogenase, complete cds	Brevibacterium saccharolyticum	066'96	13-Feb-99
	GB_OM:BTU71200	877	U71200	Bos taurus acetoin reductase mRNA, complete cds.	Bos taurus	51,659	8-Oct-97
	GB_EST2:F12685	287	F12685	HSC3DA031 normalized infant brain cDNA Homo sapiens cDNA clone c- 3da03. mRNA sequence	Homo sapiens	41,509	14-Mar-95
rxa02480 1779	GB_BA1:MTV012	70287	AL021287	Mycobacterium tuberculosis H37Rv complete genome; segment 132/162.	Mycobacterium tuberculosis	36,737	23-Jun-99

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GB_BA1:SCGG10 36734 ALU49497 GB_BA1:AP000060 347800 AP000060 GB_BA1:MTCY20G9 37218 Z77162 GB_BA1:U00018 42991 U00018 GGB_HTG2:AC008235 136017 AC008235 GB_HTG2:AC008235 136017 AC008235 GB_BA2:RSU17129 17425 U17129 GB_BA1:MTV038 16094 AL021933 GB_BA2:AF068264 3152 AF068264 GB_BA1:BACHYPTP 17057 D29985 GB_BA1:BACHUTWAPA28954 D31856 GB_BA1:BSGBGLUC 4290 Z34526
GB_BA1:SCBG10 GB_BA1:AP000060 GB_BA1:STMPGM GB_BA1:U00018 GB_BA1:U00018 GB_HTG2:AC008235 GB_HTG2:AC008235 GB_HTG2:AC008235 GB_HTG2:AC008235 GB_BA1:BACHUTWAP/GB_BA1:BA1:BACHUTWAP/GB_BA1:BACHUTWAP/GB_BA1:BACHUTWAP/GB_BA1:BACHUTWAP/GB_BA1:BACHUTWAP/GB_BA1:BACHUTWAP/GB_BA1:BACHUTWAP/GB_BA1:BACHUTWA
GB_BA1:SCBG10 GB_BA1:AP000060 GB_BA1:MTCY20G9 GB_BA1:MTCY20G9 GB_BA1:MTCY20G9 GB_HTG2:AC008235 GB_HTG2:AC008235 GB_HTG2:AC008235 GB_HTG2:AC008235 GB_BA2:RSU17129 GB_BA1:MTV038 GB_BA1:BACHYPTP GB_BA1:BACHYPTP GB_BA1:BACHYPTP GB_BA1:BACHYPTP GB_BA1:BACHYPTP GB_BA1:BACHYPTP GB_BA1:GBA1:GBA1:GBA1:GBA1:GBA1:GBA1:GBA1:G
ка02485 ка02492 840 ка02539 1641 ка02551 483

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17-Jun-98	17-Jun-98	29-Sep-99	24-Jun-99	30-OCT-1998		25-MAR-1998	17-Jun-98	10 050		29-Sep-94	17-Jun-98	10-DEC-1996	25-MAY-1995		12-Aug-99	2-Sen-99	3-Jul-99		24-Jun-99	18-MAY-1997		28-Aug-99	5-Jan-98	5-410-99	7	13-Feb-97	01-MAY-1998	
61,677	37,170	19,820	36,957	67,627	!	70,417	38,532	373 03	200	57,486	38,018	58,510	57,193		36,858	37 608	50,667		39,187	59,273		58,339	39,637	33 735	<u>.</u>	35,431	38 851	
Mycobacterium tuberculosis	Mycobacterium tuberculosis	Homo sapiens	Mycobacterium	tuberculosis s Mycobacterium	tuberculosis	• Mycobacterium	Mycobacterium	tuberculosis	tuberculosis	Mycobacterium leprae	Mycobacterium tuberculosis	Mycobacterium tuberculosis		aureofaciens	Corynebacterium	giularricum Caenorhahditis elecans	- Rattus norvegicus		Mycobacterium tuberculosis	Pseudomonas aeruginosa		Pseudomonas tolaasii	Arabidopsis thaliana	Arahidonsis thaliana		Mus musculus	Homo sapiens	
Table 4 (continued) Mycobacterium tuberculosis H37Rv complete genome; segment 16/162.	Mycobacterium tuberculosis H37Rv complete genome; segment 16/162.	Homo sapiens chromosome 21 clone LLNLc116H0124 map 21q21, *** SEQUENCING IN PROGRESS ***, in unordered pieces.	Mycobacterium tuberculosis H37Rv complete genome; segment 157/162.	tuberculosis Mycobacterium tuberculosis UDP-galactopyranose mutase (gif) gene, complete Mycobacterium	cds.	Mycobacterium tuberculosis UDP-galactopyranose mutase (gif) gene, complete Mycobacterium	cas. Mycobacterium tuberculosis H37Rv complete genome; segment 59/162.	the contract of the confidence of the contract from a few contract of Education (Contract o	Mycobackerum tubercarosis sequence non crorie y 101.	Mycobacterium leprae cosmid B1549.	Mycobacterium tuberculosis H37Rv complete genome; segment 59/162.	AD000018 Mycobacterium tuberculosis sequence from clone y151.	Streptomyces aureofaciens glycogen branching enzyme (glgB) gene, complete	cds.	Corynebacterium glutamicum yjcc gene, amtR gene and citE gene, partial.	Caanorhabditis alacans cosmid M108 completa saguence	U-R-C3-sz-h-03-0-UI.s1 UI-R-C3 Rattus norvegicus cDNA clone UI-R-C3-sz-h- Rattus norvegicus	03-0-UI 3', mRNA sequence.	Mycobacterium tuberculosis H37Rv complete genome; segment 155/162.	Pseudomonas aeruginosa (orfX), glycerol dfiffusion facilitator (glpF), glycerol	kinase (glpk), and Glp repressor (glpR) genes, complete cds, and (orfk) gene, partial cds.	Pseudomonas tolaasii glpK gene for glycerol kinase, complete cds.	20827 Lambda-PRL2 Arabidopsis thaliana cDNA clone 232B7T7, mRNA	sequence. Arabidonsis thatiana chromosome 1 RAC T17H3 semience, complete	sequence.	Mus musculus Btk locus, alpha-D-galactosidase A (Ags), ribosomal protein	(L44L), and Bruton's tyrosine kinase (Btk) genes, complete cds. Homo capiens chromosome 16, cosmid clone 363F3 (LAML), complete	sequence.
296800	Z96800	AL121632	AL022076	AF026540		U96128	Z73902	000004		U00014	Z73902	AD000018	L11647		AJ133719	746036	AI547662		121125 AL022121	<b>U49666</b>		AB015974	N65787	AC005918		U58105	AC004643	2000
38900	38900	46989	23740	1778		1200	32514	92020	3/030	36470	32514	37036	2557		1839	20073	377		121125	4495		1641	512	65830		88871	43411	<u> </u>
GB_BA1:MTCY63	GB_BA1:MTCY63	GB_HTG1:HS24H01	GB_BA1:MTV026	GB_BA2:AF026540	I	GB_BA2:MTU96128	GB_BA1:MTCY130	00 004.110000454	161 156M:15a_a5	GB_BA1:U00014	GB_BA1:MTCY130	GB_BA1:MSGY151	GB_BA1:STMGLGEN		GB_BA1:CGL133719	CB INI-CEMIOE	GB_EST29:AI547662		GB_BA1:MTV025	GB_BA1:PAU49666		GB_BA1:AB015974	GB_EST6:N65787	GB DI 2-T17H3		GB_RO:MMU58105	GR DR3.AC004643	
ка02572 668			rxa02596 1326				xa02611 1775				rxa02612 2316				rxa02621 942				rxa02640 1650				rxa02654 1008				realloger 891	14904000

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01-MAY-1998	1-Jul-98	20-MAY-1993	17-Jun-98	27-Jan-94 26-Apr-93	26-Apr-93	29-Jul-93	27-Jun-97	27-Jun-97	29-DEC-1998	27-Jun-97	15-Nov-99	24-Jun-98	19-Jun-98	27-Jul-98 24-Jun-98	12.Jul-99 27.Jul-98 20-Feb-99
41,599	40,413	40,735	36,471	38,477 57,371	57,277	57,277	50,746	36,364	37,059	42,149	37,655	99,580	38,363	39,444 98,226	60,399 36,426 99,640
Homo sapiens	Corynebacterium glutamicum	Paracoccus denitrificans e	Mycobacterium tuberculosis	Myxococcus xanthus Bacillus caldolyticus	Bacillus	Bacillus stearothermonhilus	Danio rerio	Danio rerio	A Homo sapiens	Danio rerio	Homo sapiens	Corynebacterium qlutamicum	Mycobacterium tuberculosis	Streptomyces coelicolor Corynebacterium	glutamicum Streptomyces coelicolor Streptomyces coelicolor Corynebacterium glutamicum
Table 4 (continued) Homo sapiens chromosome 16, cosmid clone 363E3 (LANL), complete	sequence.  Corynebacterium glutamicum N-acetylglutamylphosphate reductase (argC), ornithine acetyltransferase (argJ), N-acetylglutamate kinase (argB), acetylornithine transaminase (argD), ornithine carbamoyltransferase (argF), argininosuccinate synthase (argC), and arginine repressor (argR), argininosuccinate synthase (argC), and	Paracoccus dentificans NADH denydrogenase (URF4), (NQO8), (NQO9), (URF5), (URF6), (NQO10), (NQO11), (NQO12), (NQO13), and (NQO14) genes, complete cds's; biotin [acetyl-CoA carboxy] ligase (birA) gene, complete cds's;	Mycobacterium tuberculosis H37Rv complete genome; segment 101/162.	Myxococcus xanthus devR and devS genes, complete cds's. B raidolyticus lactate delydronenase (LDH) gene, complete cds.	B. stearothermophilus lct gene encoding L-lactate dehydrogenase, complete	B.stearothermophilus lct gene.	fa09d04.r1 Zebrafish ICRFzfls Danio rerio cDNA clone 11A22 5' similar to TR:G1171163 G1171163 G/T-MISMATCH BINDING PROTEIN.; mRNA sequence.	fa09d04.r1 Zebrafish ICRFzfls Danio rerio cDNA clone 11A22 5' similar to TR:G1171163 G1171163 G/T-MISMATCH BINDING PROTEIN.; mRNA sequence.	ah67d06.s1 Soares_testis_NHT Homo sapiens cDNA clone 1320683 3', mRNA Homo sapiens sequence.	199004.r1 Zebrafish ICRFzfls Danio rerio cDNA clone 11A22 5' similar to TR:G1171163 G1171163 G/T-MISMATCH BINDING PROTEIN ;; mRNA sequence.	Homo sapiens, complete sequence.	gDNA encoding glucose-6-phosphate dehydrogenase.	Mycobacterium tuberculosis H37Rv complete genome; segment 63/162.	Streptomyces coelicolor cosmid 5A7. gDNA encoding glucose-6-phosphate dehydrogenase.	Streptomyces coelicolor cosmid C22. Streptomyces coelicolor cosmid 5A7. Corynebacterium glutamicum tkt gene for transketolase, complete cds.
AC004643	AF049897	L02354	<b>Z77163</b>	L19029 M19394	M14788	A06664	AA494626	AA494626	AA758660	AA494626	150172 AC006285	E13655	Z95844	AL031107 E13655	AL096839 AL031107 AB023377
43411	9196		42861	2452	1361	1350	121	121	233	121	150172	2260	40790	40337 2260	22115 40337 2572
GB_PR3:AC004643	GB_BA2:AF049897	GB_BA1:PDENQOURF 10425	GB_BA1:MTCY339	GB_BA1:MXADEVRS	GB_BA1:BACLDHL	GB_PAT:A06664	GB_EST15:AA494626	GB_EST15:AA494626	GB_EST19:AA758660	GB_EST15:AA494626	GB_PR4:AC006285	GB_PAT:E13655	GB_BA1:MTCY493	GB_BA1:SC5A7 GB_PAT:E13655	GB_BA1:SCC22 GB_BA1:SC5A7 GB_BA1:AB023377
		ка02675 1980		px=02694 1065			rxa02729 844		rxa02730 1161			rxa02737 1665		rxa02738 1203	гха02739 2223

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04-DEC-1998	01-MAR-1994	2-Aug-99	2-Aug-99	20-Sep-99	12-Jun-98	12-Jun-98	23-Jun-99	04-DEC-1998 01-MAR-1994	26-Jun-99	8-Sep-97	08-OCT-1998	1-Feb-97	01-MAR-1994	9-Apr-97	20-Aug-98 22-Jul-99	22-Jul-99	30-Jun-99
61.573	61,573	37,105	37,105	38,728	33,116	33,116	36,379	48,401 48,401	37,128	38,889	34,321	38,072	34,462	v. 50,445	59,314 37,607	37,607	40,157
Mycobacterium leorae	Mycobacterium leprae	Drosophila melanogaster	Drosophila melanogaster	Drosophila melanogaster	Homo sapiens	Homo sapiens	Ephydatia fluviatilis	Mycobacterium leprae Mycobacterium leprae	Homo sapiens	Corynebacterium	Homo sapiens	Bacillus firmus	Mycobacterium leprae	Pseudomonas syringae pv. 50,445 syringae	Streptomyces coelicolor Homo sapiens	Homo sapiens	Mus musculus
Table 4 (continued) Mycobacterium legrae cosmid 1536.	Mycobacterium leprae cosmid B1496.	Drosophila melanogaster chromosome 2 clone BACR48110 (D505) RPCI-98 48.I.10 map 49E6-49F8 strain y; cn bw sp, *** SEQUENCING IN PROGRESS ***, 17 unordered pieces.	Drosophila melanogaster chromosome 2 clone BACR48110 (D505) RPCI-98 48.1.10 map 49E6-49F8 strain y; cn bw sp, *** SEQUENCING IN PROGRESS ***, 17 unordered pieces.	Drosophila melanogaster chromosome 2 clone BACR16P13 (D597) RPCI-98 16.P.13 map 49E-49F strain y; cn bw sp. *** SEQUENCING IN PROGRESS*** 87 unordered pieces.	Homo sapiens clone DJ1022I14, *** SEQUENCING IN PROGRESS ***, 14 unordered pieces.	Homo sapiens clone DJ1022I14, *** SEQUENCING IN PROGRESS ***, 14 unordered bleces.	Ephydatia fluviatilis mRNA for G protein a subunit 4, partial cds.	Mycobacterium feprae cosmid L536. Mycobacterium feprae cosmid B1496.	Homo sapiens clone NH0501007, *** SEQUENCING IN PROGRESS ***, 3 unordered bieces.	C.glutamicum betP gene.	HS_3136_A1_A03_MR CIT Approved Human Genomic Sperm Library D Homo sapiens genomic clone Plate=3136 Col=5 Row=A, genomic survey sequence.	Bacillus firmus dppABC operon, dipeptide transporter protein dppA gene, partial cds, and dipeptide transporter proteins dppB and dppC genes, complete cds.	Mycobacterium leprae cosmid B229.	Pseudomonas syringae pv. syringae putative dihydropteroate synthase gene, partial cds, regulatory protein MrsA (mrsA), triose phosphate isomerase (tplA), transport protein SecG (secG), tRNA-Leu, tRNA-Met, and 15 kDa protein genes, complete cds.	Streptomyces coelicolor cosmid 6G4.  Homo sapiens chromosome 17 clone 2020_K_17 map 17, *** SEQUENCING IN PROGRESS ***, 12 unordered pieces.	Homo sapiens chromosome 17 clone 2020_K_17 map 17, *** SEQUENCING IN PROGRESS ***, 12 unordered pieces.	AV117143 Mus musculus C578L/6J 10-day embryo Mus musculus cDNA clone Mus musculus 2610200J17, mRNA sequence.
Z99125	U00013	174368 AC006247	174368 AC006247	121474 AC007150	129429 AC004951	129429 AC004951	AB006546	Z99125 U00013	AC007401	X93514	AQ148714	U64514	U00020	U85643	AL031317 AC008105	AC008105	AV117143
36224	35881	174368	174368	121474	129429	129429	931	36224 35881	83657	2339	405	3837	36947	4032	41055 91421	91421	222
GB BA1:MLCL536	GB_BA1:U00013	GB_HTG2:AC006247	GB_HTG2:AC006247	GB_HTG3:AC007150	GB_HTG2:AC004951	GB_HTG2:AC004951	GB_IN1:AB006546	GB_BA1:MLCL536 GB_BA1:U00013	GB_HTG2:AC007401	GB_BA1:CGBETPGEN 2339	GB_GSS9:AQ148714	GB_BA1:BFU64514	GB_BA1:U00020	GB_BA2:PSU85643	GB_BA1:SC6G4 GB_HTG2:AC008105	GB_HTG2:AC008105	GB_EST33:AV117143
		rxa02740 1053			rxa02741 1089			ra02743 1161		xa02797 1026			rxa02803 680		rxa02821 363		

				Table 4 (continued)			
rxa02829 373	GB_HTG1:HSU9G8	48735	48735 AL008714	Homo sapiens chromosome X clone LL0XNC01-9G8, *** SEQUENCING IN PROGRESS ***, in unordered pleces.	Homo sapiens	41,595	23-Nov-99
	GB_HTG1:HSU9G8	48735	48735 AL008714	Homo sapiens chromosome X clone LL0XNC01-9G8, *** SEQUENCING IN PROGRESS ***, in unordered pieces.	Homo sapiens	41,595	23-Nov-99
	GB_PR3:HSU85B5	39550	39550 Z69724	•	Homo sapiens	41,595	23-Nov-99
אכ03216 1141	лс03216 1141 GB_HTG3:AC008184 151720 AC008184	151720	AC008184	Drosophila melanogaster chromosome 2 clone BACR04D05 (D540) RPCI-98 04.D.5 map 36E5-36F2 strain y; cn bw sp, *** SEQUENCING IN PROGRESS ***, 27 unordered pieces.	Drosophila melanogaster	39,600	2-Aug-99
	GB_EST15:AA477537 411	411	AA477537	zu36g12.r1 Soares ovary tumor NbHOT Homo sapiens cDNA clone IMAGE:740134 5' similar to contains Alu repetitive element; contains element HGR repetitive element; mRNA sequence.	Homo sapiens	37,260	9-Nov-97
	GB_EST26:Al330662	412	AI330662	fa91d08.y1 zebrafish fin day1 regeneration Danio rerio cDNA 5', mRNA sequence.	Danio rerio	37,805	28-DEC-1998
rxs03215 1038	GB_BA1:SC3F9	19830	AL023862	Streptomyces coelicolor cosmid 3F9.	Streptomyces coelicolor A3(2)	48,657	10-Feb-99
	GB_BA1:SLLINC	36270	X79146	S.lincolnensis (78-11) Lincomycin production genes.	Streptomyces lincolnensis 39,430	39,430	15-MAY-1996
	GB_HTG5:AC009660	204320			Homo sapiens	35,151	04-DEC-1999
rs03224 1288	GB_PR3:AC004076 GB_PL2:SPAC926	41322	AC004076 AL110469	Homo sapiens chromosome 19, cosmid R30217, complete sequence. S. nombe chromosome I cosmid c926.	Homo sapiens Schizosacchammyces	37,788 38 474	29-Jan-98 2-Sen-99
	GB_BA2:AE001081	11473	AE001081		pombe Archaeoglobus fulgidus	35,871	15-DEC-1997

### Exemplification

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## Example 1: Preparation of total genomic DNA of *Corynebacterium glutamicum* ATCC 13032

A culture of Corynebacterium glutamicum (ATCC 13032) was grown overnight at 30°C with vigorous shaking in BHI medium (Difco). The cells were harvested by centrifugation, the supernatant was discarded and the cells were resuspended in 5 ml buffer-I (5% of the original volume of the culture — all indicated volumes have been calculated for 100 ml of culture volume). Composition of buffer-I: 140.34 g/l sucrose, 2.46 g/l MgSO<sub>4</sub> x 7H<sub>2</sub>O, 10 ml/l KH<sub>2</sub>PO<sub>4</sub> solution (100 g/l, adjusted to pH 6.7 with KOH), 50 ml/l M12 concentrate (10 g/l (NH<sub>4</sub>)<sub>2</sub>SO<sub>4</sub>, 1 g/l NaCl, 2 g/l MgSO<sub>4</sub> x 7H<sub>2</sub>O, 0.2 g/l CaCl<sub>2</sub>, 0.5 g/l yeast extract (Difco), 10 ml/l trace-elements-mix (200 mg/l FeSO<sub>4</sub> x H<sub>2</sub>O, 10 mg/l ZnSO<sub>4</sub> x 7 H<sub>2</sub>O, 3 mg/l MnCl<sub>2</sub> x 4 H<sub>2</sub>O, 30 mg/l H<sub>3</sub>BO<sub>3</sub> 20 mg/l CoCl<sub>2</sub> x 6 H,O, 1 mg/l NiCl, x 6 H,O, 3 mg/l Na,MoO<sub>4</sub> x 2 H,O, 500 mg/l complexing agent (EDTA or critic acid), 100 ml/l vitamins-mix (0.2 mg/l biotin, 0.2 mg/l folic acid, 20 mg/l p-amino benzoic acid, 20 mg/l riboflavin, 40 mg/l ca-panthothenate, 140 mg/l nicotinic acid, 40 mg/l pyridoxole hydrochloride, 200 mg/l myo-inositol). Lysozyme was added to the suspension to a final concentration of 2.5 mg/ml. After an approximately 4 h incubation at 37°C, the cell wall was degraded and the resulting protoplasts are harvested by centrifugation. The pellet was washed once with 5 ml buffer-I and once with 5 ml TE-buffer (10 mM Tris-HCl, 1 mM EDTA, pH 8). The pellet was resuspended in 4 ml TE-buffer and 0.5 ml SDS solution (10%) and 0.5 ml NaCl solution (5 M) are added. After adding of proteinase K to a final concentration of 200 μg/ml, the suspension is incubated for ca.18 h at 37°C. The DNA was purified by extraction with phenol, phenol-chloroform-isoamylalcohol and chloroformisoamylalcohol using standard procedures. Then, the DNA was precipitated by adding 1/50 volume of 3 M sodium acetate and 2 volumes of ethanol, followed by a 30 min incubation at -20°C and a 30 min centrifugation at 12,000 rpm in a high speed centrifuge using a SS34 rotor (Sorvall). The DNA was dissolved in 1 ml TE-buffer containing 20 µg/ml RNaseA and dialysed at 4°C against 1000 ml TE-buffer for at least 3 hours. During this time, the buffer was exchanged 3 times. To aliquots of 0.4 ml of the dialysed DNA solution, 0.4 ml of 2 M LiCl and 0.8 ml of ethanol are added. After a 30

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min incubation at -20°C, the DNA was collected by centrifugation (13,000 rpm, Biofuge Fresco, Heraeus, Hanau, Germany). The DNA pellet was dissolved in TE-buffer. DNA prepared by this procedure could be used for all purposes, including southern blotting or construction of genomic libraries.

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# Example 2: Construction of genomic libraries in *Escherichia coli* of *Corynebacterium glutamicum* ATCC13032.

Using DNA prepared as described in Example 1, cosmid and plasmid libraries were constructed according to known and well established methods (see e.g., Sambrook, J. et al. (1989) "Molecular Cloning: A Laboratory Manual", Cold Spring Harbor Laboratory Press, or Ausubel, F.M. et al. (1994) "Current Protocols in Molecular Biology", John Wiley & Sons.)

Any plasmid or cosmid could be used. Of particular use were the plasmids pBR322 (Sutcliffe, J.G. (1979) *Proc. Natl. Acad. Sci. USA*, 75:3737-3741); pACYC177 (Change & Cohen (1978) *J. Bacteriol* 134:1141-1156), plasmids of the pBS series (pBSSK+, pBSSK- and others; Stratagene, LaJolla, USA), or cosmids as SuperCos1 (Stratagene, LaJolla, USA) or Lorist6 (Gibson, T.J., Rosenthal A. and Waterson, R.H. (1987) *Gene* 53:283-286. Gene libraries specifically for use in *C. glutamicum* may be constructed using plasmid pSL109 (Lee, H.-S. and A. J. Sinskey (1994) *J. Microbiol. Biotechnol.* 4: 256-263).

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### **Example 3: DNA Sequencing and Computational Functional Analysis**

Genomic libraries as described in Example 2 were used for DNA sequencing according to standard methods, in particular by the chain termination method using ABI377 sequencing machines (see *e.g.*, Fleischman, R.D. *et al.* (1995) "Whole-genome Random Sequencing and Assembly of Haemophilus Influenzae Rd., *Science*, 269:496-512). Sequencing primers with the following nucleotide sequences were used: 5'-GGAAACAGTATGACCATG-3' or 5'-GTAAAACGACGGCCAGT-3'.

## Example 4: In vivo Mutagenesis

30 In vivo mutagenesis of Corynebacterium glutamicum can be performed by passage of plasmid (or other vector) DNA through E. coli or other microorganisms (e.g. Bacillus spp. or yeasts such as Saccharomyces cerevisiae) which are impaired in their capabilities to maintain

the integrity of their genetic information. Typical mutator strains have mutations in the genes for the DNA repair system (e.g., mutHLS, mutD, mutT, etc.; for reference, see Rupp, W.D. (1996) DNA repair mechanisms, in: *Escherichia coli* and *Salmonella*, p. 2277-2294, ASM: Washington.) Such strains are well known to those of ordinary skill in the art. The use of such strains is illustrated, for example, in Greener, A. and Callahan, M. (1994) *Strategies* 7: 32-34.

# Example 5: DNA Transfer Between *Escherichia coli* and *Corynebacterium* glutamicum

Several Corvnebacterium and Brevibacterium species contain endogenous plasmids (as e.g., pHM1519 or pBL1) which replicate autonomously (for review see, e.g., Martin, J.F. et al. (1987) Biotechnology, 5:137-146). Shuttle vectors for Escherichia coli and Corynebacterium glutamicum can be readily constructed by using standard vectors for E. coli (Sambrook, J. et al. (1989), "Molecular Cloning: A Laboratory Manual", Cold Spring Harbor Laboratory Press or Ausubel, F.M. et al. (1994) "Current Protocols in Molecular Biology", John Wiley & Sons) to which a origin or replication for and a suitable marker from Corynebacterium glutamicum is added. Such origins of replication are preferably taken from endogenous plasmids isolated from Corynebacterium and Brevibacterium species. Of particular use as transformation markers for these species are genes for kanamycin resistance (such as those derived from the Tn5 or Tn903 transposons) or chloramphenicol (Winnacker, E.L. (1987) "From Genes to Clones — 20 Introduction to Gene Technology, VCH, Weinheim). There are numerous examples in the literature of the construction of a wide variety of shuttle vectors which replicate in both E. coli and C. glutamicum, and which can be used for several purposes, including gene overexpression (for reference, see e.g., Yoshihama, M. et al. (1985) J. Bacteriol. 162:591-597, Martin J.F. et al. (1987) Biotechnology, 5:137-146 and Eikmanns, B.J. et al. (1991) Gene, 25 102:93-98).

Using standard methods, it is possible to clone a gene of interest into one of the shuffle vectors described above and to introduce such a hybrid vectors into strains of *Corynebacterium glutamicum*. Transformation of *C. glutamicum* can be achieved by protoplast transformation (Kastsumata, R. et al. (1984) *J. Bacteriol*. 159306-311), electroporation (Liebl, E. et al. (1989) *FEMS Microbiol*. Letters, 53:399-303) and in cases where special vectors are used, also by conjugation (as described e.g. in Schäfer, A et al.

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(1990) J. Bacteriol. 172:1663-1666). It is also possible to transfer the shuttle vectors for C. glutamicum to E. coli by preparing plasmid DNA from C. glutamicum (using standard methods well-known in the art) and transforming it into E. coli. This transformation step can be performed using standard methods, but it is advantageous to use an Mcr-deficient E. coli strain, such as NM522 (Gough & Murray (1983) J. Mol. Biol. 166:1-19).

Genes may be overexpressed in *C. glutamicum* strains using plasmids which comprise pCG1 (U.S. Patent No. 4,617,267) or fragments thereof, and optionally the gene for kanamycin resistance from TN903 (Grindley, N.D. and Joyce, C.M. (1980) *Proc. Natl. Acad. Sci. USA* 77(12): 7176-7180). In addition, genes may be overexpressed in *C. glutamicum* strains using plasmid pSL109 (Lee, H.-S. and A. J. Sinskey (1994) *J. Microbiol. Biotechnol.* 4: 256-263).

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Aside from the use of replicative plasmids, gene overexpression can also be achieved by integration into the genome. Genomic integration in C. glutamicum or other Corynebacterium or Brevibacterium species may be accomplished by well-known methods, such as homologous recombination with genomic region(s), restriction 15 endonuclease mediated integration (REMI) (see, e.g., DE Patent 19823834), or through the use of transposons. It is also possible to modulate the activity of a gene of interest by modifying the regulatory regions (e.g., a promoter, a repressor, and/or an enhancer) by sequence modification, insertion, or deletion using site-directed methods (such as homologous recombination) or methods based on random events (such as transposon 20 mutagenesis or REMI). Nucleic acid sequences which function as transcriptional terminators may also be inserted 3' to the coding region of one or more genes of the invention; such terminators are well-known in the art and are described, for example, in Winnacker, E.L. (1987) From Genes to Clones – Introduction to Gene Technology. VCH: 25 Weinheim.

## Example 6: Assessment of the Expression of the Mutant Protein

Observations of the activity of a mutated protein in a transformed host cell rely on the fact that the mutant protein is expressed in a similar fashion and in a similar quantity to that of the wild-type protein. A useful method to ascertain the level of transcription of the mutant gene (an indicator of the amount of mRNA available for translation to the gene product) is to perform a Northern blot (for reference see, for example, Ausubel *et al.*)

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(1988) Current Protocols in Molecular Biology, Wiley: New York), in which a primer designed to bind to the gene of interest is labeled with a detectable tag (usually radioactive or chemiluminescent), such that when the total RNA of a culture of the organism is extracted, run on gel, transferred to a stable matrix and incubated with this probe, the binding and quantity of binding of the probe indicates the presence and also the quantity of mRNA for this gene. This information is evidence of the degree of transcription of the mutant gene. Total cellular RNA can be prepared from *Corynebacterium glutamicum* by several methods, all well-known in the art, such as that described in Bormann, E.R. et al. (1992) *Mol. Microbiol*. 6: 317-326.

To assess the presence or relative quantity of protein translated from this mRNA, standard techniques, such as a Western blot, may be employed (see, for example, Ausubel et al. (1988) Current Protocols in Molecular Biology, Wiley: New York). In this process, total cellular proteins are extracted, separated by gel electrophoresis, transferred to a matrix such as nitrocellulose, and incubated with a probe, such as an antibody, which specifically binds to the desired protein. This probe is generally tagged with a chemiluminescent or colorimetric label which may be readily detected. The presence and quantity of label observed indicates the presence and quantity of the desired mutant protein present in the cell.

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# 20 Example 7: Growth of Genetically Modified *Corynebacterium glutamicum* — Media and Culture Conditions

Genetically modified *Corynebacteria* are cultured in synthetic or natural growth media. A number of different growth media for Corynebacteria are both well-known and readily available (Lieb *et al.* (1989) *Appl. Microbiol. Biotechnol.*, 32:205-210; von der Osten *et al.* (1998) *Biotechnology Letters*, 11:11-16; Patent DE 4,120,867; Liebl (1992) "The Genus *Corynebacterium*, in: The Procaryotes, Volume II, Balows, A. *et al.*, eds. Springer-Verlag). These media consist of one or more carbon sources, nitrogen sources, inorganic salts, vitamins and trace elements. Preferred carbon sources are sugars, such as mono-, di-, or polysaccharides. For example, glucose, fructose, mannose, galactose, ribose, sorbose, ribulose, lactose, maltose, sucrose, raffinose, starch or cellulose serve as very good carbon sources. It is also possible to supply sugar to the media via complex compounds such as molasses or other by-products from sugar refinement. It can also be

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advantageous to supply mixtures of different carbon sources. Other possible carbon sources are alcohols and organic acids, such as methanol, ethanol, acetic acid or lactic acid. Nitrogen sources are usually organic or inorganic nitrogen compounds, or materials which contain these compounds. Exemplary nitrogen sources include ammonia gas or ammonia salts, such as NH<sub>4</sub>Cl or (NH<sub>4</sub>)<sub>2</sub>SO<sub>4</sub>, NH<sub>4</sub>OH, nitrates, urea, amino acids or complex nitrogen sources like corn steep liquor, soy bean flour, soy bean protein, yeast extract, meat extract and others.

Inorganic salt compounds which may be included in the media include the chloride-, phosphorous- or sulfate- salts of calcium, magnesium, sodium, cobalt, molybdenum, potassium, manganese, zinc, copper and iron. Chelating compounds can be added to the medium to keep the metal ions in solution. Particularly useful chelating compounds include dihydroxyphenols, like catechol or protocatechuate, or organic acids, such as citric acid. It is typical for the media to also contain other growth factors, such as vitamins or growth promoters, examples of which include biotin, riboflavin, thiamin, folic acid, nicotinic acid, pantothenate and pyridoxin. Growth factors and salts frequently originate from complex media components such as yeast extract, molasses, corn steep liquor and others. The exact composition of the media compounds depends strongly on the immediate experiment and is individually decided for each specific case. Information about media optimization is available in the textbook "Applied Microbiol. Physiology, A Practical Approach (eds. P.M. Rhodes, P.F. Stanbury, IRL Press (1997) pp. 53-73, ISBN 0 19 963577 3). It is also possible to select growth media from commercial suppliers, like standard 1 (Merck) or BHI (grain heart infusion, DIFCO) or others.

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All medium components are sterilized, either by heat (20 minutes at 1.5 bar and 121°C) or by sterile filtration. The components can either be sterilized together or, if necessary, separately. All media components can be present at the beginning of growth, or they can optionally be added continuously or batchwise.

Culture conditions are defined separately for each experiment. The temperature should be in a range between 15°C and 45°C. The temperature can be kept constant or can be altered during the experiment. The pH of the medium should be in the range of 5 to 8.5, preferably around 7.0, and can be maintained by the addition of buffers to the media. An exemplary buffer for this purpose is a potassium phosphate buffer. Synthetic buffers such as MOPS, HEPES, ACES and others can alternatively or simultaneously be used. It

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is also possible to maintain a constant culture pH through the addition of NaOH or NH<sub>4</sub>OH during growth. If complex medium components such as yeast extract are utilized, the necessity for additional buffers may be reduced, due to the fact that many complex compounds have high buffer capacities. If a fermentor is utilized for culturing the microorganisms, the pH can also be controlled using gaseous ammonia.

The incubation time is usually in a range from several hours to several days. This time is selected in order to permit the maximal amount of product to accumulate in the broth. The disclosed growth experiments can be carried out in a variety of vessels, such as microtiter plates, glass tubes, glass flasks or glass or metal fermentors of different sizes.

For screening a large number of clones, the microorganisms should be cultured in microtiter plates, glass tubes or shake flasks, either with or without baffles. Preferably 100 ml shake flasks are used, filled with 10% (by volume) of the required growth medium. The flasks should be shaken on a rotary shaker (amplitude 25 mm) using a speed-range of 100 – 300 rpm. Evaporation losses can be diminished by the maintenance of a humid atmosphere; alternatively, a mathematical correction for evaporation losses should be performed.

If genetically modified clones are tested, an unmodified control clone or a control clone containing the basic plasmid without any insert should also be tested. The medium is inoculated to an OD<sub>600</sub> of O.5 – 1.5 using cells grown on agar plates, such as CM plates (10 g/l glucose, 2,5 g/l NaCl, 2 g/l urea, 10 g/l polypeptone, 5 g/l yeast extract, 5 g/l meat extract, 22 g/l NaCl, 2 g/l urea, 10 g/l polypeptone, 5 g/l yeast extract, 5 g/l meat extract, 22 g/l agar, pH 6.8 with 2M NaOH) that had been incubated at 30°C. Inoculation of the media is accomplished by either introduction of a saline suspension of *C. glutamicum* cells from CM plates or addition of a liquid preculture of this bacterium.

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### Example 8 – In vitro Analysis of the Function of Mutant Proteins

The determination of activities and kinetic parameters of enzymes is well established in the art. Experiments to determine the activity of any given altered enzyme must be tailored to the specific activity of the wild-type enzyme, which is well within the ability of one of ordinary skill in the art. Overviews about enzymes in general, as well as specific details concerning structure, kinetics, principles, methods, applications and examples for the determination of many enzyme activities may be

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found, for example, in the following references: Dixon, M., and Webb, E.C., (1979) Enzymes. Longmans: London; Fersht, (1985) Enzyme Structure and Mechanism. Freeman: New York: Walsh, (1979) Enzymatic Reaction Mechanisms. Freeman: San Francisco: Price, N.C., Stevens, L. (1982) Fundamentals of Enzymology, Oxford Univ. Press: Oxford; Boyer, P.D., ed. (1983) The Enzymes, 3<sup>rd</sup> ed. Academic Press: New York; Bisswanger, H., (1994) Enzymkinetik, 2<sup>nd</sup> ed. VCH: Weinheim (ISBN 3527300325); Bergmeyer, H.U., Bergmeyer, J., Graßl, M., eds. (1983-1986) Methods of Enzymatic Analysis, 3<sup>rd</sup> ed., vol. I-XII, Verlag Chemie: Weinheim; and Ullmann's Encyclopedia of Industrial Chemistry (1987) vol. A9, "Enzymes". VCH: Weinheim, p. 10 352-363.

The activity of proteins which bind to DNA can be measured by several wellestablished methods, such as DNA band-shift assays (also called gel retardation assays). The effect of such proteins on the expression of other molecules can be measured using reporter gene assays (such as that described in Kolmar, H. et al. (1995) EMBO J. 14: 3895-3904 and references cited therein). Reporter gene test systems are well known and established for applications in both pro- and eukaryotic cells, using enzymes such as beta-galactosidase, green fluorescent protein, and several others.

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The determination of activity of membrane-transport proteins can be performed according to techniques such as those described in Gennis, R.B. (1989) "Pores, Channels and Transporters", in Biomembranes, Molecular Structure and Function, 20 Springer: Heidelberg, p. 85-137; 199-234; and 270-322.

## Example 9: Analysis of Impact of Mutant Protein on the Production of the Desired **Product**

The effect of the genetic modification in C. glutamicum on production of a desired compound (such as an amino acid) can be assessed by growing the modified microorganism under suitable conditions (such as those described above) and analyzing the medium and/or the cellular component for increased production of the desired product (i.e., an amino acid). Such analysis techniques are well known to one of ordinary skill in the art, and include spectroscopy, thin layer chromatography, staining 30 methods of various kinds, enzymatic and microbiological methods, and analytical chromatography such as high performance liquid chromatography (see, for example,

Ullman, Encyclopedia of Industrial Chemistry, vol. A2, p. 89-90 and p. 443-613, VCH: Weinheim (1985); Fallon, A. et al., (1987) "Applications of HPLC in Biochemistry" in: Laboratory Techniques in Biochemistry and Molecular Biology, vol. 17; Rehm et al. (1993) Biotechnology, vol. 3, Chapter III: "Product recovery and purification", page 5 469-714, VCH: Weinheim; Belter, P.A. et al. (1988) Bioseparations: downstream processing for biotechnology, John Wiley and Sons; Kennedy, J.F. and Cabral, J.M.S. (1992) Recovery processes for biological materials, John Wiley and Sons; Shaeiwitz, J.A. and Henry, J.D. (1988) Biochemical separations, in: Ulmann's Encyclopedia of Industrial Chemistry, vol. B3, Chapter 11, page 1-27, VCH: Weinheim; and Dechow, F.J. (1989) Separation and purification techniques in biotechnology, Noyes Publications.)

In addition to the measurement of the final product of fermentation, it is also possible to analyze other components of the metabolic pathways utilized for the production of the desired compound, such as intermediates and side-products, to determine the overall efficiency of production of the compound. Analysis methods include measurements of nutrient levels in the medium (e.g., sugars, hydrocarbons, nitrogen sources, phosphate, and other ions), measurements of biomass composition and growth, analysis of the production of common metabolites of biosynthetic pathways, and measurement of gasses produced during fermentation. Standard methods for these measurements are outlined in Applied Microbial Physiology, A Practical Approach, P.M. Rhodes and P.F. Stanbury, eds., IRL Press, p. 103-129; 131-163; and 165-192 (ISBN: 0199635773) and references cited therein.

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## Example 10: Purification of the Desired Product from C. glutamicum Culture

Recovery of the desired product from the C. glutamicum cells or supernatant of the above-described culture can be performed by various methods well known in the art. If the desired product is not secreted from the cells, the cells can be harvested from the culture by low-speed centrifugation, the cells can be lysed by standard techniques, such as mechanical force or sonication. The cellular debris is removed by centrifugation, and the supernatant fraction containing the soluble proteins is retained for further 30 purification of the desired compound. If the product is secreted from the C. glutamicum

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cells, then the cells are removed from the culture by low-speed centrifugation, and the supernate fraction is retained for further purification.

The supernatant fraction from either purification method is subjected to chromatography with a suitable resin, in which the desired molecule is either retained on a chromatography resin while many of the impurities in the sample are not, or where the impurities are retained by the resin while the sample is not. Such chromatography steps may be repeated as necessary, using the same or different chromatography resins. One of ordinary skill in the art would be well-versed in the selection of appropriate chromatography resins and in their most efficacious application for a particular molecule to be purified. The purified product may be concentrated by filtration or ultrafiltration, and stored at a temperature at which the stability of the product is maximized.

There are a wide array of purification methods known to the art and the preceding method of purification is not meant to be limiting. Such purification techniques are described, for example, in Bailey, J.E. & Ollis, D.F. Biochemical Engineering Fundamentals, McGraw-Hill: New York (1986).

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The identity and purity of the isolated compounds may be assessed by techniques standard in the art. These include high-performance liquid chromatography (HPLC), spectroscopic methods, staining methods, thin layer chromatography, NIRS, enzymatic assay, or microbiologically. Such analysis methods are reviewed in: Patek *et al.* (1994) *Appl. Environ. Microbiol.* 60: 133-140; Malakhova *et al.* (1996) *Biotekhnologiya* 11: 27-32; and Schmidt *et al.* (1998) *Bioprocess Engineer.* 19: 67-70. Ulmann's Encyclopedia of Industrial Chemistry, (1996) vol. A27, VCH: Weinheim, p. 89-90, p. 521-540, p. 540-547, p. 559-566, 575-581 and p. 581-587; Michal, G. (1999) Biochemical Pathways: An Atlas of Biochemistry and Molecular Biology, John Wiley and Sons; Fallon, A. *et al.* (1987) Applications of HPLC in Biochemistry in: Laboratory Techniques in Biochemistry and Molecular Biology, vol. 17.

### Example 11: Analysis of the Gene Sequences of the Invention

The comparison of sequences and determination of percent homology between two sequences are art-known techniques, and can be accomplished using a mathematical algorithm, such as the algorithm of Karlin and Altschul (1990) *Proc. Natl. Acad. Sci.* USA 87:2264-68, modified as in Karlin and Altschul (1993) *Proc. Natl. Acad. Sci.* USA

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90:5873-77. Such an algorithm is incorporated into the NBLAST and XBLAST programs (version 2.0) of Altschul, et al. (1990) J. Mol. Biol. 215:403-10. BLAST nucleotide searches can be performed with the NBLAST program, score = 100, wordlength = 12 to obtain nucleotide sequences homologous to SMP nucleic acid molecules of the invention. BLAST protein searches can be performed with the XBLAST program, score = 50, wordlength = 3 to obtain amino acid sequences homologous to SMP protein molecules of the invention. To obtain gapped alignments for comparison purposes, Gapped BLAST can be utilized as described in Altschul et al., (1997) Nucleic Acids Res. 25(17):3389-3402. When utilizing BLAST and Gapped BLAST programs, one of ordinary skill in the art will know how to optimize the parameters of the program (e.g., XBLAST and NBLAST) for the specific sequence being analyzed.

Another example of a mathematical algorithm utilized for the comparison of sequences is the algorithm of Meyers and Miller ((1988) Comput. Appl. Biosci. 4: 11-17). Such an algorithm is incorporated into the ALIGN program (version 2.0) which is part of the GCG sequence alignment software package. When utilizing the ALIGN program for comparing amino acid sequences, a PAM120 weight residue table, a gap length penalty of 12, and a gap penalty of 4 can be used. Additional algorithms for sequence analysis are known in the art, and include ADVANCE and ADAM. described in Torelli and Robotti (1994) Comput. Appl. Biosci. 10:3-5; and FASTA, described in Pearson and Lipman (1988) P.N.A.S. 85:2444-8.

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The percent homology between two amino acid sequences can also be accomplished using the GAP program in the GCG software package (available at http://www.gcg.com), using either a Blosum 62 matrix or a PAM250 matrix, and a gap weight of 12, 10, 8, 6, or 4 and a length weight of 2, 3, or 4. The percent homology between two nucleic acid sequences can be accomplished using the GAP program in the GCG software package, using standard parameters, such as a gap weight of 50 and a length weight of 3.

A comparative analysis of the gene sequences of the invention with those present in Genbank has been performed using techniques known in the art (see, e.g., Bexevanis and Ouellette, eds. (1998) Bioinformatics: A Practical Guide to the Analysis of Genes and Proteins. John Wiley and Sons: New York). The gene sequences of the invention

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were compared to genes present in Genbank in a three-step process. In a first step, a BLASTN analysis (e.g., a local alignment analysis) was performed for each of the sequences of the invention against the nucleotide sequences present in Genbank, and the top 500 hits were retained for further analysis. A subsequent FASTA search (e.g., a combined local and global alignment analysis, in which limited regions of the sequences are aligned) was performed on these 500 hits. Each gene sequence of the invention was subsequently globally aligned to each of the top three FASTA hits, using the GAP program in the GCG software package (using standard parameters). In order to obtain correct results, the length of the sequences extracted from Genbank were adjusted to the length of the query sequences by methods well-known in the art. The results of this analysis are set forth in Table 4. The resulting data is identical to that which would have been obtained had a GAP (global) analysis alone been performed on each of the genes of the invention in comparison with each of the references in Genbank, but required significantly reduced computational time as compared to such a database-wide GAP (global) analysis. Sequences of the invention for which no alignments above the cutoff values were obtained are indicated on Table 4 by the absence of alignment information. It will further be understood by one of ordinary skill in the art that the GAP alignment homology percentages set forth in Table 4 under the heading "% homology (GAP)" are listed in the European numerical format, wherein a ',' represents a decimal point. For example, a value of "40,345" in this column represents "40.345%".

### Example 12: Construction and Operation of DNA Microarrays

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The sequences of the invention may additionally be used in the construction and application of DNA microarrays (the design, methodology, and uses of DNA arrays are well known in the art, and are described, for example, in Schena, M. et al. (1995)

Science 270: 467-470; Wodicka, L. et al. (1997) Nature Biotechnology 15: 1359-1367;

DeSaizieu, A. et al. (1998) Nature Biotechnology 16: 45-48; and DeRisi, J.L. et al. (1997) Science 278: 680-686).

DNA microarrays are solid or flexible supports consisting of nitrocellulose, nylon, glass, silicone, or other materials. Nucleic acid molecules may be attached to the surface in an ordered manner. After appropriate labeling, other nucleic acids or nucleic acid mixtures can be hybridized to the immobilized nucleic acid molecules, and the label

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may be used to monitor and measure the individual signal intensities of the hybridized molecules at defined regions. This methodology allows the simultaneous quantification of the relative or absolute amount of all or selected nucleic acids in the applied nucleic acid sample or mixture. DNA microarrays, therefore, permit an analysis of the expression of multiple (as many as 6800 or more) nucleic acids in parallel (see, e.g., Schena, M. (1996) *BioEssays* 18(5): 427-431).

The sequences of the invention may be used to design oligonucleotide primers which are able to amplify defined regions of one or more *C. glutamicum* genes by a nucleic acid amplification reaction such as the polymerase chain reaction. The choice and design of the 5' or 3' oligonucleotide primers or of appropriate linkers allows the covalent attachment of the resulting PCR products to the surface of a support medium described above (and also described, for example, Schena, M. *et al.* (1995) *Science* 270: 467-470).

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Nucleic acid microarrays may also be constructed by *in situ* oligonucleotide synthesis as described by Wodicka, L. *et al.* (1997) *Nature Biotechnology* 15: 1359-1367. By photolithographic methods, precisely defined regions of the matrix are exposed to light. Protective groups which are photolabile are thereby activated and undergo nucleotide addition, whereas regions that are masked from light do not undergo any modification. Subsequent cycles of protection and light activation permit the synthesis of different oligonucleotides at defined positions. Small, defined regions of the genes of the invention may be synthesized on microarrays by solid phase oligonucleotide synthesis.

The nucleic acid molecules of the invention present in a sample or mixture of nucleotides may be hybridized to the microarrays. These nucleic acid molecules can be labeled according to standard methods. In brief, nucleic acid molecules (e.g., mRNA molecules or DNA molecules) are labeled by the incorporation of isotopically or fluorescently labeled nucleotides, e.g., during reverse transcription or DNA synthesis. Hybridization of labeled nucleic acids to microarrays is described (e.g., in Schena, M. et al. (1995) supra; Wodicka, L. et al. (1997), supra; and DeSaizieu A. et al. (1998), supra). The detection and quantification of the hybridized molecule are tailored to the specific incorporated label. Radioactive labels can be detected, for example, as

described in Schena, M. et al. (1995) supra) and fluorescent labels may be detected, for example, by the method of Shalon et al. (1996) Genome Research 6: 639-645).

The application of the sequences of the invention to DNA microarray technology, as described above, permits comparative analyses of different strains of *C. glutamicum* or other Corynebacteria. For example, studies of inter-strain variations based on individual transcript profiles and the identification of genes that are important for specific and/or desired strain properties such as pathogenicity, productivity and stress tolerance are facilitated by nucleic acid array methodologies. Also, comparisons of the profile of expression of genes of the invention during the course of a fermentation reaction are possible using nucleic acid array technology.

## Example 13: Analysis of the Dynamics of Cellular Protein Populations (Proteomics)

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The genes, compositions, and methods of the invention may be applied to study the interactions and dynamics of populations of proteins, termed 'proteomics'. Protein populations of interest include, but are not limited to, the total protein population of *C. glutamicum* (e.g., in comparison with the protein populations of other organisms), those proteins which are active under specific environmental or metabolic conditions (e.g., during fermentation, at high or low temperature, or at high or low pH), or those proteins which are active during specific phases of growth and development.

Protein populations can be analyzed by various well-known techniques, such as gel electrophoresis. Cellular proteins may be obtained, for example, by lysis or extraction, and may be separated from one another using a variety of electrophoretic techniques. Sodium dodecyl sulfate polyacrylamide gel electrophoresis (SDS-PAGE) separates proteins largely on the basis of their molecular weight. Isoelectric focusing polyacrylamide gel electrophoresis (IEF-PAGE) separates proteins by their isoelectric point (which reflects not only the amino acid sequence but also posttranslational modifications of the protein). Another, more preferred method of protein analysis is the consecutive combination of both IEF-PAGE and SDS-PAGE, known as 2-D-gel electrophoresis (described, for example, in Hermann et al. (1998) Electrophoresis 19: 3217-3221; Fountoulakis et al. (1998) Electrophoresis 19: 1193-1202; Langen et al. (1997) Electrophoresis 18: 1184-1192; Antelmann et al. (1997) Electrophoresis 18:

1451-1463). Other separation techniques may also be utilized for protein separation, such as capillary gel electrophoresis; such techniques are well known in the art.

Proteins separated by these methodologies can be visualized by standard techniques, such as by staining or labeling. Suitable stains are known in the art, and include Coomassie Brilliant Blue, silver stain, or fluorescent dyes such as Sypro Ruby (Molecular Probes). The inclusion of radioactively labeled amino acids or other protein precursors (e.g., <sup>35</sup>S-methionine, <sup>35</sup>S-cysteine, <sup>14</sup>C-labelled amino acids, <sup>15</sup>N-amino acids, <sup>15</sup>NO<sub>3</sub> or <sup>15</sup>NH<sub>4</sub><sup>+</sup> or <sup>13</sup>C-labelled amino acids) in the medium of *C. glutamicum* permits the labeling of proteins from these cells prior to their separation. Similarly, fluorescent labels may be employed. These labeled proteins can be extracted, isolated and separated according to the previously described techniques.

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Proteins visualized by these techniques can be further analyzed by measuring the amount of dye or label used. The amount of a given protein can be determined quantitatively using, for example, optical methods and can be compared to the amount of other proteins in the same gel or in other gels. Comparisons of proteins on gels can be made, for example, by optical comparison, by spectroscopy, by image scanning and analysis of gels, or through the use of photographic films and screens. Such techniques are well-known in the art.

To determine the identity of any given protein, direct sequencing or other standard techniques may be employed. For example, N- and/or C-terminal amino acid sequencing (such as Edman degradation) may be used, as may mass spectrometry (in particular MALDI or ESI techniques (see, e.g., Langen et al. (1997) Electrophoresis 18: 1184-1192)). The protein sequences provided herein can be used for the identification of C. glutamicum proteins by these techniques.

The information obtained by these methods can be used to compare patterns of protein presence, activity, or modification between different samples from various biological conditions (e.g., different organisms, time points of fermentation, media conditions, or different biotopes, among others). Data obtained from such experiments alone, or in combination with other techniques, can be used for various applications, such as to compare the behavior of various organisms in a given (e.g., metabolic) situation, to increase the productivity of strains which produce fine chemicals or to increase the efficiency of the production of fine chemicals.

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## **Equivalents**

Those of ordinary skill in the art will recognize, or will be able to ascertain using no more than routine experimentation, many equivalents to the specific embodiments of the invention described herein. Such equivalents are intended to be encompassed by the following claims.

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### What is claimed:

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- An isolated nucleic acid molecule from Corynebacterium glutamicum encoding an
   SMP protein, or a portion thereof, provided that the nucleic acid molecule does not consist of any of the F-designated genes set forth in Table 1.
  - 2. The isolated nucleic acid molecule of claim 1, wherein said nucleic acid molecule encodes an SMP protein involved in the production of a fine chemical.

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3. An isolated Corynebacterium glutamicum nucleic acid molecule selected from the group consisting of those sequences set forth as odd-numbered SEQ ID NOs of the Sequence Listing, or a portion thereof, provided that the nucleic acid molecule does not consist of any of the F-designated genes set forth in Table 1.

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4. An isolated nucleic acid molecule which encodes a polypeptide sequence selected from the group consisting of those sequences set forth as even-numbered SEQ ID NOs of the Sequence Listing,, provided that the nucleic acid molecule does not consist of any of the F-designated genes set forth in Table 1.

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- 5. An isolated nucleic acid molecule which encodes a naturally occurring allelic variant of a polypeptide selected from the group of amino acid sequences consisting of those sequences set forth in as even-numbered SEQ ID NOs of the Sequence Listing, provided that the nucleic acid molecule does not consist of any of the F-designated genes set forth in Table 1.
- 6. An isolated nucleic acid molecule comprising a nucleotide sequence which is at least 50% homologous to a nucleotide sequence selected from the group consisting of

those sequences set forth as odd-numbered SEQ ID NOs of the Sequence Listing, or

a portion thereof, provided that the nucleic acid molecule does not consist of any of the F-designated genes set forth in Table 1.

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- 7. An isolated nucleic acid molecule comprising a fragment of at least 15 nucleotides of a nucleic acid comprising a nucleotide sequence selected from the group consisting of those sequences set forth as odd-numbered SEQ ID NOs of the Sequence Listing, provided that the nucleic acid molecule does not consist of any of the F-designated genes set forth in Table 1.
  - 8. An isolated nucleic acid molecule which hybridizes to the nucleic acid molecule of any one of claims 1-7 under stringent conditions.
- 9. An isolated nucleic acid molecule comprising the nucleic acid molecule of any one of claims 1-8 or a portion thereof and a nucleotide sequence encoding a heterologous polypeptide.
  - 10. A vector comprising the nucleic acid molecule of any one of claims 1-9.

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- 11. The vector of claim 10, which is an expression vector.
- 12. A host cell transfected with the expression vector of claim 11.
- 20 13. The host cell of claim 12, wherein said cell is a microorganism.
  - 14. The host cell of claim 13, wherein said cell belongs to the genus *Corynebacterium* or *Brevibacterium*.
- 25 15. The host cell of claim 12, wherein the expression of said nucleic acid molecule results in the modulation in production of a fine chemical from said cell.
  - 16. The host cell of claim 15, wherein said fine chemical is selected from the group consisting of: organic acids, proteinogenic and nonproteinogenic amino acids, purine and pyrimidine bases, nucleosides, nucleotides, lipids, saturated and unsaturated fatty acids, diols, carbohydrates, aromatic compounds, vitamins, cofactors, polyketides, and enzymes.

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- 17. A method of producing a polypeptide comprising culturing the host cell of claim 12 in an appropriate culture medium to, thereby, produce the polypeptide.
- 5 18. An isolated SMP polypeptide from *Corynebacterium glutamicum*, or a portion thereof.
  - 19. The polypeptide of claim 18, wherein said polypeptide is involved in the production of a fine chemical.

20. An isolated polypeptide comprising an amino acid sequence selected from the group consisting of those sequences set forth as even-numbered SEQ ID NOs of the Sequence Listing, provided that the amino acid sequence is not encoded by any of

the F-designated genes set forth in Table 1.

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- 21. An isolated polypeptide comprising a naturally occurring allelic variant of a polypeptide comprising an amino acid sequence selected from the group consisting of those sequences set forth as even-numbered SEQ ID NOs of the Sequence Listing, or a portion thereof, provided that the amino acid sequence is not encoded by any of the F-designated genes set forth in Table 1.
- 22. The isolated polypeptide of any of claims 18-21, further comprising heterologous amino acid sequences.
- 25 23. An isolated polypeptide which is encoded by a nucleic acid molecule comprising a nucleotide sequence which is at least 50% homologous to a nucleic acid selected from the group consisting of those sequences set forth as odd-numbered SEQ ID NOs of the Sequence Listing, provided that the nucleic acid molecule does not consist of any of the F-designated nucleic acid molecules set forth in Table 1.

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24. An isolated polypeptide comprising an amino acid sequence which is at least 50% homologous to an amino acid sequence selected from the group consisting of those

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sequences as even-numbered SEQ ID NOs of the Sequence Listing, provided that the amino acid sequence is not encoded by any of the F-designated genes set forth in Table 1.

- 5 25. A method for producing a fine chemical, comprising culturing a cell containing a vector of claim 12 such that the fine chemical is produced.
  - 26. The method of claim 25, wherein said method further comprises the step of recovering the fine chemical from said culture.

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- 27. The method of claim 25, wherein said method further comprises the step of transfecting said cell with the vector of claim 11 to result in a cell containing said vector.
- 15 28. The method of claim 25, wherein said cell belongs to the genus *Corynebacterium* or *Brevibacterium*.
  - 29. The method of claim 25, wherein said cell is selected from the group consisting of: Corynebacterium glutamicum, Corynebacterium herculis, Corynebacterium, lilium,
- 20 Corynebacterium acetoacidophilum, Corynebacterium acetoglutamicum,
  Corynebacterium acetophilum, Corynebacterium ammoniagenes, Corynebacterium
  fujiokense, Corynebacterium nitrilophilus, Brevibacterium ammoniagenes,
  Brevibacterium butanicum, Brevibacterium divaricatum, Brevibacterium flavum,
  Brevibacterium healii, Brevibacterium ketoglutamicum, Brevibacterium
- 25 ketosoreductum, Brevibacterium lactofermentum, Brevibacterium linens, Brevibacterium paraffinolyticum, and those strains set forth in Table 3.
  - 30. The method of claim 25, wherein expression of the nucleic acid molecule from said vector results in modulation of production of said fine chemical.

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31. The method of claim 25, wherein said fine chemical is selected from the group consisting of: organic acids, proteinogenic and nonproteinogenic amino acids, purine

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and pyrimidine bases, nucleosides, nucleotides, lipids, saturated and unsaturated fatty acids, diols, carbohydrates, aromatic compounds, vitamins, cofactors, polyketides, and enzymes.

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- 5 32. The method of claim 25, wherein said fine chemical is an amino acid.
  - 33. The method of claim 32, wherein said amino acid is drawn from the group consisting of: lysine, glutamate, glutamine, alanine, aspartate, glycine, serine, threonine, methionine, cysteine, valine, leucine, isoleucine, arginine, proline, histidine, tyrosine, phenylalanine, and tryptophan.
  - 34. A method for producing a fine chemical, comprising culturing a cell whose genomic DNA has been altered by the inclusion of a nucleic acid molecule of any one of claims 1-9.

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- 35. A method for diagnosing the presence or activity of *Corynebacterium* diphtheriae in a subject, comprising detecting the presence of one or more of SEQ ID NOs 1 through 782 of the Sequence Listing in the subject, provided that the sequences are not or are not encoded by any of the F-designated sequences set forth in Table 1, thereby diagnosing the presence or activity of *Corynebacterium diphtheriae* in the subject.
- 36. A host cell comprising a nucleic acid molecule selected from the group consisting of the nucleic acid molecules set forth as odd-numbered SEQ ID NOs of the Sequence Listing, wherein the nucleic acid molecule is disrupted.
- 35. 37. A host cell comprising a nucleic acid molecule selected from the group consisting of the nucleic acid molecules set forth as odd-numbered SEQ ID NOs in the Sequence Listing, wherein the nucleic acid molecule comprises one or more nucleic acid modifications from the sequence set forth as odd-numbered SEQ ID NOs of the Sequence Listing s.

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38. A host cell comprising a nucleic acid molecule selected from the group consisting of the nucleic acid molecules set forth as odd-numbered SEQ ID NOs of the Sequence Listing, wherein the regulatory region of the nucleic acid molecule is modified
 5 relative to the wild-type regulatory region of the molecule.

## SEQUENCE LISTING

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cgc gca cgc gat act gaa gat ttg gtt gca cag gct gcc tcc aaa ttc 1 Arg Ala Arg Asp Thr Glu Asp Leu Val Ala Gln Ala Ala Ser Lys Phe 10 15 20	.63
att gag gtt gtt gaa gca gca act gcc aat aat ggc acc gca cag gta Ile Glu Val Val Glu Ala Ala Thr Ala Asn Asn Gly Thr Ala Gln Val 25 30 35	211
gtg ctc acc ggt ggt ggc gcc ggc atc aag ttg ctg gaa aag ctc agc 2 Val Leu Thr Gly Gly Gly Ala Gly Ile Lys Leu Glu Lys Leu Ser 40 45 50	259
gtt gat gcg gct gac ctt gcc tgg gat cgc att cat gtg ttc ttc ggc 3 Val Asp Ala Ala Asp Leu Ala Trp Asp Arg Ile His Val Phe Phe Gly 55 60 65	307
gat gag cgc aat gtc cct gtc agt gat tct gag tcc aat gag ggc cag Asp Glu Arg Asn Val Pro Val Ser Asp Ser Glu Ser Asn Glu Gly Gln 70 80 85	355
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cac gga tat ggt ctc ggc gac gta gat ctt gca gag gca gcc cgc gct 4 His Gly Tyr Gly Leu Gly Asp Val Asp Leu Ala Glu Ala Ala Arg Ala 105 110 115	51
tac gaa gct gtg ttg gat gaa ttc gca cca aac ggc ttt gat ctt cac 4 Tyr Glu Ala Val Leu Asp Glu Phe Ala Pro Asn Gly Phe Asp Leu His 120 125 130	99
ctg ctc ggc atg ggt ggc gaa ggc cat atc aac tcc ctg ttc cct cac 5	47

1

Бей	Leu 135	Gly	Met	Gly	Gly	Glu 140	Gly	His	Ile	Asn	Ser 145	Leu	Phe	Pro	His	
						tcc Ser										595
						gag Glu										643
		-	_	-		tgg Trp	_	-	-					_	-	691
						gtc Val										739
						tct Ser 220										787
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2

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25

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25 30 35

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40 45 50

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Thr Val Asn Asp Pro Gly Ser Leu Gly Ile Val Leu Gly Gly Ser Gly
55 60 65

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cg Ar 23	c tgc g Cys 0	gaa Glu	ggc Gly	ctg Leu	atc Ile 235	atg Met	cgc Arg	aac Asn	ctg Leu	atc Ile 240	acc Thr	ggc Gly	gag Glu	ctc Leu	acc Thr 245	835
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	g tac l Tyr															931
	g cgt t Arg															979
	g ttc n Phe 295															1027
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	t aag o Lys														Asp	1123
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	c cca l Pro															1219
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Pro Ala Gly Val Pro Thr Lys Asp Met Trp Glu Tyr Gln Lys Asp His  $35 \hspace{1cm} 40 \hspace{1cm} 45$ 

Met Asn Leu Val Ser Pro Leu Asn Arg Arg Lys Phe Arg Val Leu Val 50 55 60

Val Gly Thr Gly Leu Ser Gly Gly Ala Ala Ala Ala Ala Leu Gly Glu 65 70 75 80

Leu Gly Tyr Asp Val Lys Ala Phe Thr Tyr His Asp Ala Pro Arg Arg 85 90 95

Ala His Ser Ile Ala Ala Gln Gly Gly Val Asn Ser Ala Arg Gly Lys 100 105 110

Lys Val Asp Asn Asp Gly Ala Tyr Arg His Val Lys Asp Thr Val Lys 115 120 125

Gly Gly Asp Tyr Arg Gly Arg Glu Ser Asp Cys Trp Arg Leu Ala Val 130 135 140

Glu Ser Val Arg Val Ile Asp His Met Asn Ala Ile Gly Ala Pro Phe 145 150 155 160

Ala Arg Glu Tyr Gly Gly Ala Leu Ala Thr Arg Ser Phe Gly Gly Val 165 170 175

Gln Val Ser Arg Thr Tyr Tyr Thr Arg Gly Gln Thr Gly Gln Gln Leu 180 185 190

Gln Phe Ser Thr Ala Ser Ala Leu Gln Arg Gln Ile His Leu Gly Ser 195 200 205

Val Glu Ile Phe Thr His Asn Glu Met Val Asp Val Ile Val Thr Glu 210 215 220

Arg Asn Gly Glu Lys Arg Cys Glu Gly Leu Ile Met Arg Asn Leu Ile 225 230 235 240

Thr Gly Glu Leu Thr Ala His Thr Gly His Ala Val Ile Leu Ala Thr 245 250 255

Gly Gly Tyr Gly Asn Val Tyr His Met Ser Thr Leu Ala Lys Asn Ser 260 265 270

Asn Ala Ser Ala Ile Met Arg Ala Tyr Glu Ala Gly Ala Tyr Phe Ala 275 280 285

Ser Pro Ser Phe Ile Gln Phe His Pro Thr Gly Leu Pro Val Asn Ser 290 295 300

Thr Trp Gln Ser Lys Thr Ile Leu Met Ser Glu Ser Leu Arg Asn Asp 310 315 Gly Arg Ile Trp Ser Pro Lys Glu Pro Asn Asp Asn Arg Asp Pro Asn Thr Ile Pro Glu Asp Glu Arg Asp Tyr Phe Leu Glu Arg Arg Tyr Pro Ala Phe Gly Asn Leu Val Pro Arg Asp Val Ala Ser Arg Ala Ile Ser Gln Gln Ile Asn Ala Gly Leu Gly Val Gly Pro Leu Asn Asn Ala Ala Tyr Lew Asp Phe Arg Asp Ala Thr Glu Arg Leu Gly Gln Asp Thr Ile Arg Glu Arg Tyr Ser Asn Leu Phe Thr Met Tyr Glu Glu Ala Ile Gly Glu Asp Pro Tyr Ser Ser Pro Met Arg Ile Ala Pro Thr Cys His Phe Thr Met Gly Gly Leu Trp Thr Asp Phe Asn Glu Met Thr Ser Leu Pro 440 Gly Leu Phe Cys Ala Gly Glu Ala Ser Trp Thr Tyr His Gly Ala Asn 460 Arg Leu Gly Ala Asn Ser Leu Leu Ser Ala Ser Val Asp Gly Trp Phe 470 Thr Leu Pro Phe Thr Ile Pro Asn Tyr Leu Gly Pro Leu Leu Gly Ser Glu Arg Leu Ser Glu Asp Ala Pro Glu Ala Gln Ala Ala Ile Ala Arg Ala Gln Ala Arg Ile Asp Arg Leu Met Gly Asn Arg Pro Glu Trp Val Gly Asp Asn Val His Gly Pro Glu Tyr Tyr His Arg Gln Leu Gly Asp Ile Leu Tyr Phe Ser Cys Gly Val Ser Arg Asn Val Glu Asp Leu Gln Asp Gly Ile Asn Lys Ile Arg Ala Leu Arg Asp Asp Phe Trp Lys Asn 570 Met Arg Ile Thr Gly Ser Thr Asp Glu Met Asn Gln Val Leu Glu Tyr Ala Ala Arg Val Ala Asp Tyr Ile Asp Leu Gly Glu Leu Met Cys Val 600 Asp Ala Leu Asp Arg Asp Glu Ser Cys Gly Ala His Phe Arg Asp Asp 610 His Leu Ser Glu Asp Gly Glu Ala Gln Arg Asp Asp Gln Asn Trp Cys

630

625

635

640

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145	150	155	160
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ggc cca ttg ctt gg Gly Pro Leu Leu Gl 180	c tcc gag cgt ctg ; y Ser Glu Arg Leu ; 185	tca gag gat gca cca gaa Ser Glu Asp Ala Pro Glu 190	gca 576 Ala
		cgc att gac cgc ctc atg Arg Ile Asp Arg Leu Met 205	
aac cgc cca gag tg Asn Arg Pro Glu Tr 210	g gtc ggt gac aac o p Val Gly Asp Asn ' 215	gtt cac gga cct gag tac Val His Gly Pro Glu Tyr 220	tac 672 Tyr
cac cgc cag ctt gg His Arg Gln Leu Gl 225	c gat atc ctg tac y Asp Ile Leu Tyr 230	ttc tcc tgt ggc gtt tcc Phe Ser Cys Gly Val Ser 235	cga 720 Arg 240
aac gta gaa gac ct Asn Val Glu Asp Le 24	u Gln Asp Gly Ile 7	aac aag atc cgt gcc ctc Asn Lys Ile Arg Ala Leu 250 255	cgc 768 Arg
gat gac ttc tgg aa Asp Asp Phe Trp Ly 260	g aac atg cgc atc s Asn Met Arg Ile o	acc ggc agc acc gat gag Thr Gly Ser Thr Asp Glu 270	atg 816 Met
		gta gcc gac tac atc gac Val Ala Asp Tyr Ile Asp 285	
		gac cgc gac gag tcc tgt Asp Arg Asp Glu Ser Cys 300	
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gac gac caa aac to Asp Asp Gln Asn Tr 32	p Cys Phe Val Ser	gca tgg gaa cca ggc gag Ala Trp Glu Pro Gly Glu 330 335	aat 1008 Asn
		ctg ttc ttc gaa tct gtc Leu Phe Phe Glu Ser Val 350	
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Leu	Glu	Arg 35	Arg	Tyr	Pro	Ala	Phe 40	Gly	Asn	Leu	Val	Pro 45	Arg	Asp	Val
Ala	Ser 50	Arg	Ala	Ile	Ser	Gln 55	Gln	Ile	Asn	Ala	Gly 60	Leu	Gly	Val	Gly
Pro 65	Leu	Asn	Asn	Ala	Ala 70	Tyr	Leu	Asp	Phe	Arg 75	Asp	Ala	Thr	Glu	Arg 80
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Tyr	Glu	Glu	Ala 100	Ile	Gly	Glu	Asp	Pro 105	Tyr	Ser	Ser	Pro	Met 110	Arg	Ile
Ala	Pro	Thr 115	Cys	His	Phe	Thr	Met 120	Gly	Gly	Leu	Trp	Thr 125	Asp	Phe	Asn
Glu	Met 130	Thr	Ser	Leu	Pro	Gly 135	Leu	Phe	Cys	Ala	Gly 140	Glu	Ala	Ser	Trp
Thr 145	Tyr	His	Gly	Ala	Asn 150	Arg	Leu	Gly	Ala	Asn 155	Ser	Leu	Leu	Ser	Ala 160
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Gln	Ala	Ala 195	Ile	Ala	Arg	Ala	Gln 200	Ala	Arg	Ile	Asp	Arg 205	Leu	Met	Gly
Asn	Arg 210	Pro	Glu	Trp	Val	Gly 215	Asp	Asn	Val	His	Gly 220	Pro	Glu	Tyr	Tyr
His 225	Arg	Gln	Leu	Gly	Asp 230	Ile	Leu	Tyr	Phe	Ser 235	Суѕ	Gly	Val	Ser	Arg 240
Asn	Val	Glu	Asp	Leu 245	Gln	Asp	Gly	Ile	Asn 250	Lys	Ile	Arg	Ala	Leu 255	Arg
Asp	Asp	Phe	Trp 260	Lys	Asn	Met	Arg	Ile 265	Thr	Gly	Ser	Thr	Asp 270	Glu	Met
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Gly	Glu 290	Leu	Met	Cys	Val	Asp 295	Ala	Leu	Asp	Arg	Asp 300	Glu	Ser	Cys	Gly
Ala 305	His	Phe	Arg	Asp	Asp 310	His	Leu	Ser	Glu	Asp 315	Gly	Glu	Ala	Gln	Arg 320
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Met Thr Ile Asn Val

1 5

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ctg gct gct ctt gat gct gca tgc gct gtt cag gcc gag tgg gct agg 307 Leu Ala Ala Leu Asp Ala Ala Cys Ala Val Gln Ala Glu Trp Ala Arg 55 60 . 65

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gta gca gaa cgt gca gaa gag ttc gcc acc ctc atg acc ttg gaa atg 403 Val Ala Glu Arg Ala Glu Glu Phe Ala Thr Leu Met Thr Leu Glu Met

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				gca Ala											643
				ctt Leu											691
-		-	_	ggt Gly			-		-				-		739
 -		-		gcg Ala				_		_	_	-	_	-	787
				ttc Phe 235											835
	_	-	_	gat Asp		-	_	-			-	_			883
				att Ile											931
				ggt Gly											979
-	_		_	ttc Phe		-		-		-	-	_	_		1027
Arg	Arg	Phe	Āla	gcc Ala 315	Arg	Leu	Ğlu	Glu	Gln	Val					1075
				acc Thr											1123
				ctt Leu											1171
				aag Lys											1219
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						aac Asn										1363
						tca Ser										1411
						gtc Val										1459
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Val	Glu	Asn 35	Pro	Ala	Thr	Gly	Glu 40	Thr	Ile	Ala	Thr	Leu 45	Ala	Ser	Ala	
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Ala 65	Glu	Trp	Ala	Arg	Met 70	Pro	Ala	Arg	Glu	Arg 75	Ser	Asn	Ile	Leu	Arg 80	
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Met Thr Leu Glu Met Gly Lys Pro Leu Ala Glu Ala Arg Gly Glu Val

Thr Tyr Gly Asn Glu Phe Leu Arg Trp Phe Ser Glu Glu Ala Val Arg 115 120 125

Leu Tyr Gly Arg Tyr Gly Thr Thr Pro Glu Gly Asn Leu Arg Met Leu 130  $$135\$ 

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	gtt Val															691
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tcc Ser	ctc Leu 215	tac Tyr	ggc Gly	gag Glu	tgc Cys	gca Ala 220	gat Asp	gtc Val	tgc Cys	ccc Pro	gca Ala 225	ggc Gly	atc Ile	cca Pro	ctg Leu	787
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<211> 249

<212> PRT

<213> Corynebacterium glutamicum

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Lys Glu Pro Phe Ala Phe Ala Ser Asp Cys Arg Glu Gly Ile Cys Gly 50 60

Thr Cys Gly Leu Leu Val Asn Gly Arg Pro His Gly Ala Asp Gln Asn 65 70 75 80

Lys Pro Ala Cys Ala Gln Arg Leu Val Ser Tyr Lys Glu Gly Asp Thr \$85\$ 90 95

Leu Lys Ile Glu Pro Leu Arg Ser Ala Ala Tyr Pro Val Ile Lys Asp 100 105 110

Met Val Val Asp Arg Ser Ala Leu Asp Arg Val Met Glu Gln Gly Gly 115 120 125

Tyr Val Thr Ile Asn Ala Gly Thr Ala Pro Asp Ala Asp Thr Leu His 130 135 140

145	ASII	птэ	GIU	Int	150	GIU	rea	Ala	Leu	155	HIS	АТА	Ala	Cys	11e 160	
Gly	Cys	Gly	Ala	Cys 165	Val	Ala	Ala	Cys	Pro 170	Asn	Gly	Ala	Ala	His 175	Leu	
Phe	Thr	Gly	Ala 180	Lys	Leu	Val	His	Leu 185	Ser	Leu	Leu	Pro	Leu 190	Gly	Lys	
Glu	Glu	Arg 195	Gly	Leu	Arg	Ala	Arg 200	Lys	Met	Val	Asp	Glu 205	Met	Glu	Thr	
Asn	Phe 210	Gly	His	Cys	Ser	Leu 215	Tyr	Gly	Glu	Cys	Ala 220	Asp	Val	Cys	Pro	
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	1> CI 2> (:	101).	(15	507)												
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<400 according to the second test of the second tes	cgt Arg ctg Leu ggt	gta t  gta t  att  Ile  tgg  Trp  cgt  Arg  40  gct	gag Glu cag Gln 25 ggt Gly	cac His 10 gca Ala ctg Leu	gac Asp cag Gln	acc Thr acc Thr tcc Ser	atg Met cag Gln gca Ala 45	ggt Gly cgc Arg 30 cag Gln	gaa Glu 15 gct Ala atc Ile	gtg Val gtt Val cgc Arg	atg Met 1 aag Lys gag Glu gca Ala	acc Thr gtt Val aac Asn atg Met 50	gag Glu cca Pro ttc Phe 35 ggt Gly	cag Gln gca Ala 20 cct Pro ctg Leu	gaa Glu 5 aag Lys atc Ile ctg Leu	115 163 211
<400 according to the get Ala tet Ser aag Lys	cgt Arg ctg Leu ggt Gly gca Ala 55	gta to the state of the state o	gag Glu cag Gln 25 ggt Gly tgt Cys	cac His 10 gca Ala ctg Leu gcg Ala	gac Asp cag Gln gaa Glu	acc Thr acc Thr tcc Ser gta Val 60	atg Met cag Gln gca Ala 45 aac Asn	ggt Gly cgc Arg 30 cag Gln aag Lys	gaa Glu 15 gct Ala atc Ile gac Asp	gtg Val gtt Val cgc Arg tcc Ser	atg Met 1 aag Lys gag Glu gca Ala ggt Gly 65	acc Thr gtt Val aac Asn atg Met 50 gcg Ala	gag Glu cca Pro ttc Phe 35 ggt Gly ctg Leu	cag Gln gca Ala 20 cct Pro ctg Leu gat Asp	gaa Glu 5 aag Lys atc Ile ctg Leu gca Ala	<ul><li>115</li><li>163</li><li>211</li><li>259</li></ul>

100 90 95 451 act tee tee aac atg aac ace aat gag gtt ate get tee ate geg aag Thr Ser Ser Asn Met Asn Thr Asn Glu Val Ile Ala Ser Ile Ala Lys 110 105 499 qct aac qqc qtt qaq gtt cac cca aat gac cac gtc aac atg ggt cag Ala Asn Gly Val Glu Val His Pro Asn Asp His Val Asn Met Gly Gln 125 120 tec tec aat gac ace tte cet act gea act cac gtt get gea ace gaa 547 Ser Ser Asn Asp Thr Phe Pro Thr Ala Thr His Val Ala Ala Thr Glu 135 140 595 gct qct qtc aat gac ctc atc cca ggc ctg aag gtt ctg cac gag tct Ala Ala Val Asn Asp Leu Ile Pro Gly Leu Lys Val Leu His Glu Ser 155 ttg qcq aag aag gct aac gag tgg tct gag gtt gtt aag tcc ggc cgc 643 Leu Ala Lys Lys Ala Asn Glu Trp Ser Glu Val Val Lys Ser Gly Arg 170 acc cac ctg atg gac gct gtt cca gta acc ctg ggc cag gag ttc ggt Thr His Leu Met Asp Ala Val Pro Val Thr Leu Gly Gln Glu Phe Gly 190 qqc tac qct cqc caq atc caq ctc qqc atc gag cqc gtt gag gct act 739 Gly Tyr Ala Arg Gln Ile Gln Leu Gly Ile Glu Arg Val Glu Ala Thr 200 205 210 ctt cct cgc ctt ggt gag ctg gct att ggt ggc acc gct gct ggt acc · 787 Leu Pro Arg Leu Gly Glu Leu Ala Ile Gly Gly Thr Ala Ala Gly Thr 220 215 ggt atc aac acc tcc gct gat ttc ggc ggc aag gtt gtt gct gaa ctg 835 Gly Ile Asn Thr Ser Ala Asp Phe Gly Gly Lys Val Val Ala Glu Leu 230 235 atc aac ttg acc gac gtc aag gag ctc aag gaa gct gag aac cac ttc Ile Asn Leu Thr Asp Val Lys Glu Leu Lys Glu Ala Glu Asn His Phe qaq qct cag qct qca cqc gac qct ctt gtt gag ttc tcc ggc gca atg Glu Ala Gln Ala Arg Asp Ala Leu Val Glu Phe Ser Gly Ala Met 265 979 cgc gtt atc gct gtc tcc ttg tac aag atc gct aac gat atc cgc ctc Arg Val Ile Ala Val Ser Leu Tyr Lys Ile Ala Asn Asp Ile Arg Leu 280 285 atg ggc tcc ggc cca ctg acc ggt ctt ggc gag atc cgt ctc cca gac 1027 Met Gly Ser Gly Pro Leu Thr Gly Leu Gly Glu Ile Arg Leu Pro Asp 295 300 ctq caq cca ggt tcc tcc atc atg cca ggc aag gtc aac cca gtt ctc 1075 Leu Gln Pro Gly Ser Ser Ile Met Pro Gly Lys Val Asn Pro Val Leu 310 315 tgt gag acc gct acc cag gtt tcc gct cag gtt atc ggc aat gac gca 1123 Cys Glu Thr Ala Thr Gln Val Ser Ala Gln Val Ile Gly Asn Asp Ala

335

gct Ala	gtt Val	gcg Ala	ttc Phe 345	tcc Ser	ggc Gly	acc Thr	cag Gln	ggc Gly 350	cag Gln	ttc Phe	gag Glu	ctc Leu	aac Asn 355	gtg Val	ttc Phe	1171
atc Ile	cca Pro	gtg Val 360	atg Met	gct Ala	cgc Arg	aac Asn	gtg Val 365	ctt Leu	gag Glu	tcc Ser	gct Ala	cgc Arg 370	ctg Leu	ctg Leu	gct Ala	1219
							acc Thr									1267
aac Asn 390	gag Glu	gca Ala	cac His	atg Met	aag Lys 395	gag Glu	ctc Leu	gct Ala	gag Glu	tct Ser 400	tca Ser	cct Pro	tcc Ser	atc Ile	gtt Val 405	1315
							ggc Gly									1363
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Gln Thr Gly Ser Gly Thr Ser Ser Asn Met Asn Thr Asn Glu Val Ile

100 105 110 Ala Ser Ile Ala Lys Ala Asn Gly Val Glu Val His Pro Asn Asp His 120 Val Asn Met Gly Gln Ser Ser Asn Asp Thr Phe Pro Thr Ala Thr His 135 Val Ala Ala Thr Glu Ala Ala Val Asn Asp Leu Ile Pro Gly Leu Lys 150 Val Leu His Glu Ser Leu Ala Lys Lys Ala Asn Glu Trp Ser Glu Val Val Lys Ser Gly Arg Thr His Leu Met Asp Ala Val Pro Val Thr Leu Gly Gln Glu Phe Gly Gly Tyr Ala Arg Gln Ile Gln Leu Gly Ile Glu Arg Val Glu Ala Thr Leu Pro Arg Leu Gly Glu Leu Ala Ile Gly Gly Thr Ala Ala Gly Thr Gly Ile Asn Thr Ser Ala Asp Phe Gly Gly Lys 230 235 Val Val Ala Glu Leu Ile Asn Leu Thr Asp Val Lys Glu Leu Lys Glu Ala Glu Asn His Phe Glu Ala Gln Ala Ala Arg Asp Ala Leu Val Glu 260 265 Phe Ser Gly Ala Met Arg Val Ile Ala Val Ser Leu Tyr Lys Ile Ala 280 Asn Asp Ile Arg Leu Met Gly Ser Gly Pro Leu Thr Gly Leu Gly Glu 295 Ile Arg Leu Pro Asp Leu Gln Pro Gly Ser Ser Ile Met Pro Gly Lys Val Asn Pro Val Leu Cys Glu Thr Ala Thr Gln Val Ser Ala Gln Val Ile Gly Asn Asp Ala Ala Val Ala Phe Ser Gly Thr Gln Gly Gln Phe Glu Leu Asn Val Phe Ile Pro Val Met Ala Arg Asn Val Leu Glu Ser Ala Arg Leu Leu Ala Asn Thr Ser Arg Val Phe Ala Thr Arg Leu Val 375 Asp Gly Ile Glu Pro Asn Glu Ala His Met Lys Glu Leu Ala Glu Ser 385 390 Ser Pro Ser Ile Val Thr Pro Leu Asn Ser Ala Ile Gly Tyr Glu Ala 405 410

Ala Ala Lys Val Ala Lys Thr Ala Leu Ala Glu Gly Lys Thr Ile Arg

425

Gln Thr Val Ile Asp Leu Gly Leu Val Asp Gly Glu Lys Leu Thr Glu 435 Glu Glu Leu Asp Lys Arg Leu Asp Val Leu Ala Met Ala His Thr Glu Arg Glu Asn Lys Phe <210> 19 <211> 1164 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(1141) <223> RXA00517 <400> 19 qqtcttaqaa ccaqcqtqca ctgatqqcqa ttaaaqqqqq ttqcqcctat acctattqct 60 ggtatacatt tcggtatacc taaaccgaat tgagggattc atg cca gaa gtc act Met Pro Glu Val Thr gtc aac gcc caa caa ctc act gtt ctc tgc aca gac atc ctc acc aaa Val Asn Ala Gln Gln Leu Thr Val Leu Cys Thr Asp Ile Leu Thr Lys 10 act qqa qta cct qca qca qac qcc cat ctt qtc ggt gat agt ttg gtg 211 Thr Gly Val Pro Ala Ala Asp Ala His Leu Val Gly Asp Ser Leu Val 25 30 cag get gat ett tgg ggt cac eec tee cac ggt gtg ett ega etg eet 259 Gln Ala Asp Leu Trp Gly His Pro Ser His Gly Val Leu Arg Leu Pro 40 tgg tat gtg cgc aga ctc cac agt ggc gcg atg act aca cat gca cac 307 Trp Tyr Val Arg Arg Leu His Ser Gly Ala Met Thr Thr His Ala His 55 355 gtg gag gtt ctc aat gat ttg ggt gcc gtg ttg gcg ttg gat gga cac Val Glu Val Leu Asn Asp Leu Gly Ala Val Leu Ala Leu Asp Gly His 70 aat gga atc ggc caa gtt tta gct gat cat gct cgt aaa gaa gca gtg 403 Asn Gly Ile Gly Gln Val Leu Ala Asp His Ala Arg Lys Glu Ala Val 90 act agg gca atg atg ttc ggc atc ggt gcg gtg tcg gtg cgc aac tcc Thr Arg Ala Met Met Phe Gly Ile Gly Ala Val Ser Val Arg Asn Ser 105 499 aat cat ttt gga act gcc atg tac tac acc cgg aaa gcg gca gcg caa Asn His Phe Gly Thr Ala Met Tyr Tyr Thr Arg Lys Ala Ala Ala Gln 120 gga tgt gtt tcc att ctc acc acc aat gca tct ccg gcg atg gcg ccc 547

Gly	Cys 135	Val	Ser	Ile	Leu	Thr 140	Thr	Asn	Ala	Ser	Pro 145	Ala	Met	Ala	Pro	
					aaa Lys 155											595
					acg Thr											643
					atc Ile											691
				-	atc Ile	-	-			-			-	-		739
-		-			ggt Gly	-	-			-	-					787
					atg Met 235											835
	_		-		aag Lys	-			_		_					883
					ttg Leu											931
					gat Asp											979
			_		gca Ala											1027
					gcg Ala 315											1075
			Lys		tgg Trp											1123
	-	-		cac His	cgt Arg	tgat	ctgo	egc g	gttaa	aacct	g go	cc				1164

<sup>&</sup>lt;210> 20

<sup>&</sup>lt;211> 347

<sup>&</sup>lt;212> PRT

<sup>&</sup>lt;213> Corynebacterium glutamicum

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315

Ala His Gly Ile Ser Leu Pro Glu Lys Thr Trp Met Glu Leu Gln Glu 325 330 335

Leu Ala Ile Glu Asn His Val Val Thr His Arg

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29

145

595

140

cca gat gtt cca gca tcc cgc ttc aac gca atg atg cgc ctt gat cac

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					cag Gln											643
-					gtg Val	•						_		_		691
					gca Ala											739
					tat Tyr											787
					att Ile 235											835
-	-				att Ile	-		-	_	_		_				883
					gcg Ala											931
					gtc Val											979
					ggc Gly											1027
	_			_	cag Gln 315	-	_	_	-		_		-	-	_	1075
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Glu Val Phe Gly Thr Asp Thr Pro Val Glu Leu Lys Leu Leu Glu Ile

<sup>&</sup>lt;211> 328

<sup>&</sup>lt;212> PRT

<sup>&</sup>lt;213> Corynebacterium glutamicum

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Arg Glu Ala Val Arg Asp Leu Leu 325

295

Phe Gln Arg Ala Arg Ile Asp Ala Asn Ala Gln Glu Leu Gln Ala Glu

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<211> 1092

<212> DNA

<213> Corynebacterium glutamicum

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					aat Asn											787
-	~			_	ctc Leu 235	-	_	_	_	_		-				835
	_		-		atc Ile		-	-	-		_			-		883
					ctg Leu			-	-	-			-	-	-	931
					tcc Ser											979
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<210> 24

<211> 323

<212> PRT

<213> Corynebacterium glutamicum

<400> 24

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Ser Leu Ser Ala Pro Ser Pro Arg Thr Thr Gln Ala Met Glu Gln Gly 35 40 45

Ile Phe Asp Leu Val Glu Gln Leu Lys Ala Glu Tyr Pro Val Gly Ala 50 55 60

Val Gly Leu Ala Val Ala Gly Phe Leu Asp Pro Glu Cys Glu Val Val 65 70 75 80

Arg Phe Ala Pro His Leu Pro Trp Arg Asp Glu Pro Val Arg Glu Lys
85 90 95

Leu Glu Asn Leu Leu Gly Leu Pro Val Arg Leu Glu His Asp Ala Asn 100 105 110

Ser Ala Ala Trp Gly Glu His Arg Phe Gly Ala Ala Gln Gly Ala Asp Asn Trp Val Leu Leu Ala Leu Gly Thr Gly Ile Gly Ala Ala Leu Ile Glu Lys Gly Glu Ile Tyr Arg Gly Ala Tyr Gly Thr Ala Pro Glu Phe Gly His Leu Arg Val Val Arg Gly Gly Arg Ala Cys Ala Cys Gly Lys Glu Gly Cys Leu Glu Arg Tyr Cys Ser Gly Thr Ala Leu Val Tyr Thr Ala Arg Glu Leu Ala Ser His Gly Ser Phe Arg Asn Ser Gly Leu Phe Asp Lys Ile Lys Ala Asp Pro Asn Ser Ile Asn Gly Lys Thr Ile Thr Ala Ala Arg Gln Glu Asp Pro Leu Ala Leu Ala Val Leu Glu Asp 235 Phe Ser Glu Trp Leu Gly Glu Thr Leu Ala Ile Ile Ala Asp Val Leu 245 250 Asp Pro Gly Met Ile Ile Gly Gly Gly Leu Ser Asn Ala Ala Asp Leu Tyr Leu Asp Arg Ser Val Asn His Tyr Ser Thr Arg Ile Val Gly 280 285 Ala Gly Tyr Arg Pro Leu Ala Arg Val Ala Thr Ala Gln Leu Gly Ala 295 Asp Ala Gly Met Ile Gly Val Ala Asp Leu Ala Arg Arg Ser Val Val 310 315 Glu Ala Asn <210> 25 <211> 1785 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(1762) <223> RXA01814 <400> 25 tgttaagcca ccctactccg tgaattttgc cgtatctcgt gcgcacaatt gcttttgagg 60 qqaaqatgaa gaqaaagtat tggtgtttta aggagcaaac atg gca cat gaa cgc Met Ala His Glu Arg

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gct ttc as Ala Phe As 55			Leu Ala					307
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cgc gat ac		Leu Ser					Glu	403
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ggc atc at Gly Ile II 135		_		_				547
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aca Thr	ttg Leu	gac Asp 280	acc Thr	gac Asp	ggc Gly	aag Lys	atc Ile 285	cgc Arg	atg Met	gac Asp	tgc Cys	tcc Ser 290	agc Ser	cca Pro	cac His	979
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490 500 495 gga gga cta aaa gtg acc acc gaa aac gcc tgg ttc gca gca cgc cca 1651 Gly Gly Leu Lys Val Thr Thr Glu Asn Ala Trp Phe Ala Ala Arg Pro 505 tcc ggc acc gaa gac aag tac aag atc tac gca gaa tcc ttc aag ggc 1699 Ser Gly Thr Glu Asp Lys Tyr Lys Ile Tyr Ala Glu Ser Phe Lys Gly 520 gaa gag cac ctc gcc cag gtt cag aag gaa gcc caa gcg ttg gtc agc 1747 Glu Glu His Leu Ala Gln Val Gln Lys Glu Ala Gln Ala Leu Val Ser 535 gaa gta ctc gga cag taaaactgcg gacttgctga caa 1785 Glu Val Leu Gly Gln 550 <210> 26 <211> 554 <212> PRT <213> Corynebacterium glutamicum <400> 26 Met Ala His Glu Arg Ala Gly Gln Leu Ala Gln Pro Glu Asp Leu Ile Asp Val Ala Glu Leu Val Thr Ala Tyr Phe Thr Arg Lys Pro Asp Val 25 Asn Asn Pro Asp Gln Gln Val Ala Phe Gly Thr Ser Gly His Arg Gly 35 Phe Ala Leu Asp Ser Ala Phe Asn Glu Asp His Ile Leu Ala Thr Thr Gln Ala Ile Val Asp Tyr Arg Asn Gln Gln Pro Lys Asn Trp Val Gly Pro Leu Phe Ile Gly Arg Asp Thr His Ala Leu Ser Glu Pro Ala Met Ile Ser Ala Leu Glu Val Leu Ile Ala Asn Asp Val Glu Val Leu Val Asp Ala Asp Gly Arg Tyr Thr Pro Thr Pro Ala Val Ser His Ala Ile 120 Leu Arg His Asn Asp Gly Ile Ile Leu Gly Thr Ala Gly Pro Ser Arg 130 Pro Tyr Ala Asp Gly Ile Val Ile Thr Pro Ser His Asn Pro Pro Arg 150 155

Asp Gly Gly Phe Lys Tyr Asn Pro Ala Asn Gly Gly Pro Ala Asp Thr

Asp Ala Thr Asp Trp Ile Ala Asn Arg Ala Asn Asp Ile Leu Arg Gly

185

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Asn 225	Val	Val	Asn	Ile	Asp 230	Ala	Ile	Arg	Glu	Ala 235	Gly	Val	Arg	Ile	Gly 240
Ala	Asp	Pro	Met	Gly 245	Gly	Ala	Ser	Val	Asp 250	Tyr	Trp	Gly	Ala	Ile 255	Ala
Glu	Thr	His	Gly 260	Leu	Asn	Leu	Thr	Val 265	Val	Asn	Pro	His	Val 270	Asp	Ser
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Gln 465	Lys	Ala	Ile	Leu	Lys 470	Ala	Leu	Ser	Pro	Glu 475	Gln	Val	Thr	Ala	Thr 480
Glu	Leu	Ala	Gly	Glu 485	Ala	Ile	Thr	Ala	Lys 490	Leu	Thr	Glu	Ala	Pro 495	Gly
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528

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atc aac atg gac tgc ggt tcc acc cac att gat cag gcg cag ccg cca

150

Ile Asn Met Asp Cys Gly Ser Thr His Ile Asp Gln Ala Gln Pro Pro 170 qtc ttg aag cac ggt gct gac ctt gga ctc gcg cat gac ggt gat gct 576 Val Leu Lys His Gly Ala Asp Leu Gly Leu Ala His Asp Gly Asp Ala 185 624 qac cqt tqt ttq qct qtg aac aag gat gcc aac ctt gtt gat ggt gac Asp Arg Cys Leu Ala Val Asn Lys Asp Ala Asn Leu Val Asp Gly Asp 200 caa atc atg gcg ctg tta gcc att gcg atg aaa taaaacggcg agctgcgcaa 677 Gln Ile Met Ala Leu Leu Ala Ile Ala Met Lys 215 680 gaa <210> 28 <211> 219 <212> PRT <213> Corynebacterium glutamicum <400> 28 Val Ser Gly Glu Met Leu Ala Ala Leu Ser Ala Gly Met Ala Ser Gln Gly Val Asp Val Ile Arg Val Gly Val Ile Pro Thr Pro Ala Val 20 25 Ala Phe Leu Thr Asp Asp Tyr Gly Ala Asp Met Gly Val Met Ile Ser Ala Ser His Asn Pro Met Pro Asp Asn Gly Ile Lys Phe Phe Ser Ala Gly Gly His Lys Leu Pro Asp His Val Glu Asp Glu Ile Glu Arg Val Met Asp Ser Leu Pro Ala Glu Gly Pro Thr Gly His Gly Val Gly Arg Val Ile Glu Glu Ala Thr Asp Ala Gln Asp Arg Tyr Leu Glu His Leu 105 Lys Glu Ala Val Pro Thr Ser Leu Glu Gly Ile Lys Ile Val Val Asp Ala Ala Asn Gly Ala Ala Ser Val Val Ala Pro Thr Ala Tyr Glu Ala 135 Ala Gly Ala Thr Val Ile Ala Ile His Asn Lys Pro Asp Ser Tyr Asn 145 Ile Asn Met Asp Cys Gly Ser Thr His Ile Asp Gln Ala Gln Pro Pro 170 Val Leu Lys His Gly Ala Asp Leu Gly Leu Ala His Asp Gly Asp Ala Asp Arg Cys Leu Ala Val Asn Lys Asp Ala Asn Leu Val Asp Gly Asp

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<213> Corynebacterium glutamicum

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Phe	Leu	Thr 35	Asp	Asp	Tyr	Gly	Ala 40	Asp	Met	Gly	Val	Met 45	Ile	Ser	Ala	
Ser	His 50	Asn	Pro	Met	Pro	Asp 55	Asn	Gly	Ile	Lys	Phe 60	Phe	Ser	Ala	Gly	
Gly 65	His	Lys	Leu	Pro	Asp 70	His	Val	Glu	Asp	Glu 75	Ile	Glu	Arg	Val	Met 80	
Asp	Ser	Leu	Pro	Ala 85	Glu	Gly	Pro	Thr	Gly 90	His	Gly	Val	Gly	Arg 95	Val	
Ile	Glu	Glu	Ala 100	Thr	Asp	Ala	Gln	Asp 105	Arg	Tyr	Leu	Glu	His 110	Leu	Lys	
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	ctt Leu															163
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	cct Pro 55															307
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gtg Val	acg Thr	ttg Leu	ctc Leu 105	ccc Pro	acg Thr	cct Pro	agc Ser	cct Pro 110	acg Thr	ccg Pro	ttg Leu	att Ile	ccg Pro 115	tgg Trp	ttg Leu	451
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cag Gln 150	ctt Leu	tat Tyr	tct Ser	gaa Glu	ctg Leu 155	gag Glu	cct Pro	gag Glu	ctt Leu	gag Glu 160	gcg Ala	cat His	atc Ile	aat Asn	gct Ala 165	595
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	cag Gln															739
	gtg Val 215															787
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WO 01/00844	PCT/IB00/00943

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gtg ccg gat aag gac Val Pro Asp Lys Asp 390			
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Pro	Val 50	Pro	His	Leu	Val	Pro 55	Glu	Asp	Glu	Thr	Gly 60	Ile	Gly	Arg	Ala
Leu 65	Tyr	Pro	Gln	Asp	Gly 70	Pro	Leu	Arg	Val	Val 75	Val	Gly	Tyr	Asp	Ala 80
Arg	Tyr	Gly	Ser	His 85	Thr	Phe	Ala	Ala	Thr 90	Thr	Ala	Glu	Val	Phe 95	Ala
Gly	Ala	Gly	Phe 100	Glu	Val	Thr	Leu	Leu 105	Pro	Thr	Pro	Ser	Pro 110	Thr	Pro
Leu	Ile	Pro 115	Trp	Leu	Val	Asn	Lys 120	His	Gly	Leu	Asp	Ala 125	Gly	Val	Glr
Ile	Thr 130	Ala	Ser	His	Asn	Gly 135	Ala	Ala	Asp	Asn	Gly 140	Tyr	Lys	Val	Phe
Leu 145	Ser	Asn	Gly	Arg	Gln 150	Leu	Tyr	Ser	Glu	Leu 155	Glu	Pro	Glu	Leu	Glu 160
Ala	His	Ile	Asn	Ala 165	Val	Glu	Asp	Pro	Ile 170	Arg	Val	Pro	Arg	Val 175	Thr
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Arg 225	Ala	Met	Alà	Asn	Ala 230	Phe	Gln	Phe	Ala	Gly 235	Phe	Pro	His	Thr	His 240
Gly	Val	Lys	Ala	Gln 245	Gln	Tyr	Pro	Asp	Pro 250	Thr	Phe	Pro	Thr	Val 255	Ala
Phe	Pro	Asn	Pro 260	Glu	Glu	Pro	Ser	Ala 265	Ile	Glu	Leu	Leu	Leu 270	Glu	Arç
Ala	Lys	Glu 275	Lys	Asn	Ala	Asp	Ile 280	Leu	Phe	Ala	Leu	Asp 285	Pro	Asp	Ala
Asp	Arg 290	Cys	Ala	Val	Gly	Ile 295	Arg	Thr	Ala	Asp	Gly 300	Gly	His	Arg	Met
Leu 305	Ser	Gly	Asp	Glu	Val 310	Gly	Thr	Leu	Leu	Ala 315	Thr	Arg	Leu	Val	Pro 320

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						gat Asp										499
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						gtt Val										691
						cgg Arg										739
						cat His 220										787
						ttt Phe										835
cag	tat	cct	gat	ccc	acc	ttc	ccc	act	gtg	gcg	ttc	ccc	aat	ccg	gaa	883

Gln Tyr	Pro Asp	Pro Thr 250	Phe	Pro	Thr	Val 255	Ala	Phe	Pro	Asn	Pro 260	Glu	
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	att ttg Ile Leu 280												979
	cgt acc Arg Thr												1027
	aca ctt Thr Leu		Thr										1075
	cgt ccc Arg Pro												1123
	atc gcc Ile Ala 345												1171
	aaa aat Lys Asn 360												1219
	gct tat Ala Tyr												1267
	gat aag Asp Lys		/ Ile				_		_		_		1315
-	gaa ctg Glu Leu		_		-	-	-	-					1363
	tat cgc Tyr Arg 425	-						_			-		1411
	agc agt Ser Ser 440												1459
	gaa ctc Glu Leu												1507
	ggc att Gly Ile		His										1555
-	ggt cga Gly Arg	-						-	_			-	1603

490 495 500

gaa gtt ggt cag gcc agc tcc cat gat gaa gca gct cag ttg ttg cat 1651 Glu Val Gly Gln Ala Ser Ser His Asp Glu Ala Ala Gln Leu Leu His 505 510 515

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Pro Val Pro His Leu Val Pro Glu Asp Glu Thr Gly Ile Gly Arg Ala 50 55 60

Leu Tyr Pro Gln Asp Gly Pro Leu Arg Val Val Val Gly Tyr Asp Ala 65 70 75. 80

Arg Tyr Gly Ser His Thr Phe Ala Ala Thr Thr Ala Glu Val Phe Ala  $85 \\ 90 \\ 95$ 

Gly Ala Gly Phe Glu Val Thr Leu Leu Pro Thr Pro Ser Pro Thr Pro 100 105 110

Leu Ile Pro Trp Leu Val Asn Lys His Gly Leu Asp Ala Gly Val Gln 115 120 125

Ile Thr Ala Ser His Asn Gly Ala Ala Asp Asn Gly Tyr Lys Val Phe 130 135 140

Leu Ser Asn Gly Arg Gln Leu Tyr Ser Glu Leu Glu Pro Glu Leu Glu 145 150 155 160

Ala His Ile Asn Ala Val Glu Asp Pro Ile Arg Val Pro Arg Val Thr
165 170 175

Val Arg Pro Thr Ala Asp Gln Leu Arg Arg Tyr Val Asp Glu Met Val 180 185 190

Ser Leu Val Thr Pro Asp Gln Ala Asp Leu Leu Arg Val Asn Ser Glu 195 200 205

Arg Gly Asn Leu Arg Val Val Tyr Thr Ala Leu His Gly Val Gly 210 215 220

Arg Ala Met Ala Asn Ala Phe Gln Phe Ala Gly Phe Pro His Thr His 225 230 235 240

Gly Val Lys Ala Gln Gln Tyr Pro Asp Pro Thr Phe Pro Thr Val Ala Phe Pro Asn Pro Glu Glu Pro Ser Ala Ile Glu Leu Leu Glu Arg Ala Lys Glu Lys Asn Ala Asp Ile Leu Phe Ala Leu Asp Pro Asp Ala 280 Asp Arg Cys Ala Val Gly Ile Arg Thr Ala Asp Gly Gly His Arg Met Leu Ser Gly Asp Glu Val Gly Thr Leu Leu Ala Thr Arg Leu Val Pro 315 Glu Tyr Ser Gly Glu Gly Pro Arg Pro Val Val Ala Thr Thr Val Val Ser Ser Gln Leu Leu Gly Ile Ile Ala Glu Asp Lys Gly Trp Asp Tyr Ser Glu Thr Leu Thr Gly Phe Lys Asn Leu Ser Arg Ala Ala Asp Gly 360 Leu Asp Gly Pro Leu Ala Phe Ala Tyr Glu Glu Ala Val Gly Thr Cys 375 Pro Val Pro Asp Val Val Pro Asp Lys Asp Gly Ile Ser Thr Ala Leu 390 395 Phe Met Ala Ser Trp Ala Ala Glu Leu Lys Ala Gln Gly Ala Ser Leu 405 Gln Gln Lys Leu Asn Glu Leu Tyr Arg Arg Tyr Gly Tyr Phe Ala Ser Ser Gln Ile Ala Val Arg Thr Ser Ser Pro Arg Glu Leu Val Asp His Trp Ile Ala His Pro Gln Gln Glu Leu Ile Gly Val Ser Val Thr Pro His Ile Leu Pro Glu Lys Gln Gly Ile Ala Leu His Gly Gln Val Gly His Val His Ile Arg Ala Ile Gly Arg Val Ser Gly Thr Glu Ala Lys 490 Ala Lys Leu Tyr Leu Glu Val Gly Gln Ala Ser Ser His Asp Glu Ala Ala Gln Leu Leu His Gln Leu Glu Asp Glu Val Gln Ser Trp Leu Ser 520 525

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ttg aag att gct atg gat gaa gcc gga att aca ctg cgt acc acc aag
Leu Lys Ile Ala Met Asp Glu Ala Gly Ile Thr Leu Arg Thr Thr Lys
gta gga gac cgc tac gtg ctg gaa gac ctc aat gca ggt gga ttc agc
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Val Gly Asp Arg Tyr Val Leu Glu Asp Leu Asn Ala Gly Gly Phe Ser
                             40
ctg ggc ggc gag caa tct ggc cac att gtt ctt cca gat cat ggc acc
                                                                   192
Leu Gly Gly Glu Gln Ser Gly His Ile Val Leu Pro Asp His Gly Thr
                         55
                                                                   240
act ggc gat gga act ttg act ggt ctt tcc atc atg gcg cgc atg gct
Thr Gly Asp Gly Thr Leu Thr Gly Leu Ser Ile Met Ala Arg Met Ala
gaa acc gga aag tcc ttg ggc gag ttg gca caa gct atg acg gtg ctg
                                                                   288
Glu Thr Gly Lys Ser Leu Gly Glu Leu Ala Gln Ala Met Thr Val Leu
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cca cag gtt ctg atc aat gtg cca gtt tcg gat aag tcc acc atc gtg
Pro Gln Val Leu Ile Asn Val Pro Val Ser Asp Lys Ser Thr Ile Val
                                105
age cae cea age gtt gtg get geg ate geg gaa gea gaa get gag ttg
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Ser His Pro Ser Val Val Ala Ala Ile Ala Glu Ala Glu Ala Glu Leu
                            120
gge gee ace ggt ege gtt ett ett egt get tet gge ace gaa gag ett
                                                                   432
Gly Ala Thr Gly Arg Val Leu Leu Arg Ala Ser Gly Thr Glu Glu Leu
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ttc cgc gtg atg gtt gag gct gga gac aag gaa caa gct cgt cgt atc
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Phe Arg Val Met Val Glu Ala Gly Asp Lys Glu Gln Ala Arg Arg Ile
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Arg	Leu	Met 40	Arg	Ser	Glu	Gly	Glu 45	Thr	Thr	Val	Ala	Ile 50	Gly	His	Asp	
atg Met	cgt Arg 55	gat Asp	tcc Ser	tcc Ser	cct Pro	gaa Glu 60	ttg Leu	gcc Ala	aag Lys	gcg Ala	ttt Phe 65	gcc Ala	gat Asp	ggc Gly	gtg Val	307
act Thr 70	gca Ala	cag Gln	ggt Gly	ttg Leu	gat Asp 75	gtt Val	gtt Val	cat His	ttg Leu	gga Gly 80	ctg Leu	act Thr	tct Ser	act Thr	gat Asp 85	355
	ctg Leu															403
act Thr	gcg Ala	tcg Ser	cat His 105	aac Asn	ccc Pro	gct Ala	gag Glu	tac Tyr 110	aac Asn	ggc Gly	atc Ile	aag Lys	ttg Leu 115	tgt Cys	cgt Arg	451
gcg Ala	ggt Gly	gct Ala 120	cgt Arg	ccg Pro	gtc Val	ggt Gly	cag Gln 125	gat Asp	tct Ser	ggt Gly	ttg Leu	gcc Ala 130	aac Asn	atc Ile	att Ile	499
gat Asp	gat Asp 135	ctg Leu	gtt Val	gag Glu	ggt Gly	gtt Val 140	cca Pro	gcg Ala	ttt Phe	gat Asp	ggt Gly 145	gag Glu	tca Ser	ggt Gly	tcg Ser	547
	tct Ser															595
ctt Leu	gtt Val	gat Asp	ctg Leu	aag Lys 170	aac Asn	atc Ile	cgc Arg	cċg Pro	atg Met 175	aag Lys	gtt Val	gct Ala	gtg Val	gat Asp 180	gcg Ala	643
gca Ala	aac Asn	ggc Gly	atg Met 185	ggt Gly	ggg Gly	ttc Phe	act Thr	gtc Val 190	cct Pro	gag Glu	gta Val	ttc Phe	aag Lys 195	ggt Gly	ctg Leu	691
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	cat His 215															787
aag Lys 230	ttt Phe	acc Thr	gta Val	gag Glu	acc Thr 235	gga Gly	tct Ser	gat Asp	atc Ile	ggt Gly 240	ttg Leu	gcg Ala	ttc Phe	gac Asp	ggc Gly 245	835
gat Asp	gcg Ala	gat Asp	cgt Arg	tgc Cys 250	ttc Phe	gtg Val	gtc Val	gat Asp	gag Glu 255	aag Lys	ggc Gly	cag Gln	cca Pro	gtc Val 260	agc Ser	883
cct Pro	tcg Ser	gcg Ala	atc Ile 265	tgt Cys	gcg Ala	atc Ile	gta Val	gcg Ala 270	gag Glu	cgt Arg	tac Tyr	ttg Leu	gag Glu 275	aag Lys	ctt Leu	931

280 285 290 1027 qaq qtq att gct gaa aac ggt ggc act gcg gtg cgt act cgc gtg ggt Glu Val Ile Ala Glu Asn Gly Gly Thr Ala Val Arg Thr Arg Val Gly 300 cac tee tte ate aaq geg aag atg gea gag ace ggt geg gee ttt ggt 1075 His Ser Phe Ile Lys Ala Lys Met Ala Glu Thr Gly Ala Ala Phe Gly 315 ggc gag cac tot gcg cac tac tac ttc act gag ttc ttc aat gcg gac 1123 Gly Glu His Ser Ala His Tyr Tyr Phe Thr Glu Phe Phe Asn Ala Asp tcc ggc att ttg gct gcg atg cac gtg ctg gct gcg ctg gga agc cag 1171 Ser Gly Ile Leu Ala Ala Met His Val Leu Ala Ala Leu Gly Ser Gln 345 gac cag cca ctc agt gag atg atg gct agg tat aac cgg tac gtt gct 1219 Asp Gln Pro Leu Ser Glu Met Met Ala Arg Tyr Asn Arg Tyr Val Ala 360 365 tca ggc gag ttg aac tcc cgt ttg gct aat gca gag gcg cag caa gag Ser Gly Glu Leu Asn Ser Arg Leu Ala Asn Ala Glu Ala Gln Glu 380 385 375 ege ace cag get gtg ete gat geg tte get gat ege ace gag tee gtg 1315 Arg Thr Gln Ala Val Leu Asp Ala Phe Ala Asp Arg Thr Glu Ser Val 390 395 400 gac acc ctt gac ggc gtg act gtg gaa ctc aag gac acc tcc gcg tgg 1363 Asp Thr Leu Asp Gly Val Thr Val Glu Leu Lys Asp Thr Ser Ala Trp 420 410 ttc aac gtg cgt gcg tcc aac acc gag ccg ctg ctt cgc ctc aat gtt 1411 Phe Asn Val Arg Ala Ser Asn Thr Glu Pro Leu Leu Arg Leu Asn Val 430 425 qaa qct qca tcq aaq gaa gaa gtc gat gcg ttg gta gcg gag att cta 1459 Glu Ala Ala Ser Lys Glu Glu Val Asp Ala Leu Val Ala Glu Ile Leu 440 445 ggg att atc cgc gca taatcccatt ttccggcggg cat 1497 Gly Ile Ile Arg Ala 455 <210> 38 <211> 458 <212> PRT <213> Corynebacterium glutamicum Met Arq Thr Arq Glu Ser Val Thr Ala Val Ile Lys Ala Tyr Asp Val Arg Gly Val Val Gly Val Asp Ile Asp Ala Asp Phe Ile Ser Glu Thr Gly Ala Ala Phe Gly Arg Leu Met Arg Ser Glu Gly Glu Thr Thr Val

40

45

Ala Ile Gly His Asp Met Arg Asp Ser Ser Pro Glu Leu Ala Lys Ala Phe Ala Asp Gly Val Thr Ala Gln Gly Leu Asp Val Val His Leu Gly Leu Thr Ser Thr Asp Glu Leu Tyr Phe Ala Ser Gly Thr Leu Lys Cys Ala Gly Ala Met Phe Thr Ala Ser His Asn Pro Ala Glu Tyr Asn Gly 105 Ile Lys Leu Cys Arg Ala Gly Ala Arg Pro Val Gly Gln Asp Ser Gly Leu Ala Asn Ile Ile Asp Asp Leu Val Glu Gly Val Pro Ala Phe Asp Gly Glu Ser Gly Ser Val Ser Glu Gln Asp Leu Leu Ser Ala Tyr Ala Glu Tyr Leu Asn Glu Leu Val Asp Leu Lys Asn Ile Arg Pro Met Lys Val Ala Val Asp Ala Ala Asn Gly Met Gly Gly Phe Thr Val Pro Glu Val Phe Lys Gly Leu Pro Leu Asp Val Ala Pro Leu Tyr Phe Glu Leu Asp Gly Asn Phe Pro Asn His Glu Ala Asn Pro Leu Glu Pro Ala Asn Leu Val Asp Leu Gln Lys Phe Thr Val Glu Thr Gly Ser Asp Ile Gly 230 235 Leu Ala Phe Asp Gly Asp Ala Asp Arg Cys Phe Val Val Asp Glu Lys 250 245 Gly Gln Pro Val Ser Pro Ser Ala Ile Cys Ala Ile Val Ala Glu Arg Tyr Leu Glu Lys Leu Pro Gly Ser Thr Ile Ile His Asn Leu Ile Thr Ser Lys Ala Val Pro Glu Val Ile Ala Glu Asn Gly Gly Thr Ala Val Arq Thr Arg Val Gly His Ser Phe Ile Lys Ala Lys Met Ala Glu Thr Gly Ala Ala Phe Gly Gly Glu His Ser Ala His Tyr Tyr Phe Thr Glu Phe Phe Asn Ala Asp Ser Gly Ile Leu Ala Ala Met His Val Leu Ala Ala Leu Gly Ser Gln Asp Gln Pro Leu Ser Glu Met Met Ala Arg Tyr 355 360

Asn Arg Tyr Val Ala Ser Gly Glu Leu Asn Ser Arg Leu Ala Asn Ala Glu Ala Gln Gln Glu Arg Thr Gln Ala Val Leu Asp Ala Phe Ala Asp 390 Arg Thr Glu Ser Val Asp Thr Leu Asp Gly Val Thr Val Glu Leu Lys Asp Thr Ser Ala Trp Phe Asn Val Arg Ala Ser Asn Thr Glu Pro Leu 425 Leu Arg Leu Asn Val Glu Ala Ala Ser Lys Glu Glu Val Asp Ala Leu Val Ala Glu Ile Leu Gly Ile Ile Arg Ala <210> 39 <211> 994 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(994) <223> FRXA01365 <400> 39 cctgatcagg acgaatcata aggtttgcta ttcggattgg atcctttggc aggggtagga 60 ttgcaagcgt tattttgttc cctaacccct tcgaggattt atg cgt acc cgt gaa Met Arg Thr Arg Glu tct gtc aca gct gta att aag gcg tat gac gtc cgt ggt gtt gtt ggt Ser Val Thr Ala Val Ile Lys Ala Tyr Asp Val Arg Gly Val Val Gly 10 gtc gat att gat gct gat ttc att tct gag act ggc gct gcc ttt ggt 211 Val Asp Ile Asp Ala Asp Phe Ile Ser Glu Thr Gly Ala Ala Phe Gly 25 259 cgg ctc atg cgt agt gag ggt gaa acc acc gtt gct att ggc cat gac Arg Leu Met Arg Ser Glu Gly Glu Thr Thr Val Ala Ile Gly His Asp 40 45 307 atg cgt gat tcc tcc cct gaa ttg gcc aag gcg ttt gcc gat ggc gtg Met Arg Asp Ser Ser Pro Glu Leu Ala Lys Ala Phe Ala Asp Gly Val 55 act gca cag ggt ttg gat gtt gtt cat ttg gga ctg act tct act gat 355 Thr Ala Gln Gly Leu Asp Val Val His Leu Gly Leu Thr Ser Thr Asp 70 gag ctg tac ttt gcg tcc gga acc ttg aag tgt gct ggt gcg atg ttt 403 Glu Leu Tyr Phe Ala Ser Gly Thr Leu Lys Cys Ala Gly Ala Met Phe 90 95 100 451 act gcg tcg cat aac ccc gct gag tac aac ggc atc aag ttg tgt cgt

56

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gat gat ctg Asp Asp Leu 135							
gtt tct gag Val Ser Glu 150	Gln Asp L						
ctt gtt gat Leu Val Asp							
gca aac ggc Ala Asn Gly	atg ggt g Met Gly G 185	ggg ttc a Sly Phe T	act gtc Thr Val 190	cct gag Pro Glu	gta ttc Val Phe	aag ggt Lys Gly 195	ctg 691 Leu
cca ctt gat Pro Leu Asp 200		ro Leu T					
aac cat gag Asn His Glu 215							
aag ttt acc Lys Phe Thr 230	Val Glu T						
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Gly Ala Ala Phe Gly Arg Leu Met Arg Ser Glu Gly Glu Thr Thr Val 35 40 45

Ala Ile Gly His Asp Met Arg Asp Ser Ser Pro Glu Leu Ala Lys Ala 50 60

Phe Ala Asp Gly Val Thr Ala Gln Gly Leu Asp Val Val His Leu Gly 65 70 75 80

Leu Thr Ser Thr Asp Glu Leu Tyr Phe Ala Ser Gly Thr Leu Lys Cys 85 90 95

Ala Gly Ala Met Phe Thr Ala Ser His Asn Pro Ala Glu Tyr Asn Gly 100 105 110

Ile Lys Leu Cys Arg Ala Gly Ala Arg Pro Val Gly Gln Asp Ser Gly 115 120 125

Leu Ala Asn Ile Ile Asp Asp Leu Val Glu Gly Val Pro Ala Phe Asp 130 135 140

Gly Glu Ser Gly Ser Val Ser Glu Gln Asp Leu Leu Ser Ala Tyr Ala 145 150 155 160

Glu Tyr Leu Asn Glu Leu Val Asp Leu Lys Asn Ile Arg Pro Met Lys 165 170 175

Val Ala Val Asp Ala Ala Asn Gly Met Gly Gly Phe Thr Val Pro Glu 180 185 190

Val Phe Lys Gly Leu Pro Leu Asp Val Ala Pro Leu Tyr Phe Glu Leu 195 200 205

Asp Gly Asn Phe Pro Asn His Glu Ala Asn Pro Leu Glu Pro Ala Asn 210 215 220

Leu Val Asp Leu Gln Lys Phe Thr Val Glu Thr Gly Ser Asp Ile Gly 225 230 235 240

Leu Ala Phe Asp Gly Asp Ala Asp Arg Cys Phe Val Val Asp Glu Lys 245 250 255

Gly Gln Pro Val Ser Pro Ser Ala Ile Cys Ala Ile Val Ala Glu Arg 260 265 270

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Ser Lys Ala Val Pro Glu Val Ile Ala Glu 290 295

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200 205 210 tcc aac gct cgt gca gct cgt gct tgg ctg gta gag aag ctc ggt gaa 787 Ser Asn Ala Arg Ala Arg Ala Trp Leu Val Glu Lys Leu Gly Glu 220 qaq qct qtc qcq aaq cac ttc qtc qca gtg tcc acc aat gct gaa aag 835 Glu Ala Val Ala Lys His Phe Val Ala Val Ser Thr Asn Ala Glu Lys 235 gtc gca gag ttc ggt atc gac acg gac aac atg ttc ggc ttc tgg gac 883 Val Ala Glu Phe Gly Ile Asp Thr Asp Asn Met Phe Gly Phe Trp Asp 250 tgg gtc gga ggt cgt tac tcc gtg gac tcc gca gtt ggt ctt tcc ctc 931 Trp Val Gly Gly Arg Tyr Ser Val Asp Ser Ala Val Gly Leu Ser Leu 270 atq gca gtg atc ggc cct cgc gac ttc atg cgt ttc ctc ggt gga ttc 979 Met Ala Val Ile Gly Pro Arg Asp Phe Met Arg Phe Leu Gly Gly Phe 280 285 cac gcg atg gat gaa cac ttc cgc acc acc aag ttc gaa gag aac gtt 1027 His Ala Met Asp Glu His Phe Arg Thr Thr Lys Phe Glu Glu Asn Val 300 295 cca atc ttq atg gct ctg ctc ggt gtc tgg tac tcc gat ttc tat ggt 1075 Pro Ile Leu Met Ala Leu Leu Gly Val Trp Tyr Ser Asp Phe Tyr Gly 315 320 gca gaa acc cac gct gtc cta cct tat tcc gag gat ctc agc cgt ttt 1123 Ala Glu Thr His Ala Val Leu Pro Tyr Ser Glu Asp Leu Ser Arg Phe 330 340 gct gct tac ctc cag cag ctg acc atg gaa tca aat ggc aag tca gtc 1171 Ala Ala Tyr Leu Gln Gln Leu Thr Met Glu Ser Asn Gly Lys Ser Val 345 cac ege gae gge tee eet gtt tee aet gge aet gge gaa att tae tgg His Arg Asp Gly Ser Pro Val Ser Thr Gly Thr Gly Glu Ile Tyr Trp 360 ggt gag cct ggc aca aat ggc cag cac gct ttc ttc cag ctg atc cac 1267 Gly Glu Pro Gly Thr Asn Gly Gln His Ala Phe Phe Gln Leu Ile His 375 380 cag ggc act cgc ctt qtt cca gct gat ttc att ggt ttc gct cgt cca 1315 Gln Gly Thr Arg Leu Val Pro Ala Asp Phe Ile Gly Phe Ala Arg Pro 400 390 395 aag cag gat ctt cct gcc ggt gag cgc acc atg cat gac ctt ttg atg 1363 Lys Gln Asp Leu Pro Ala Gly Glu Arg Thr Met His Asp Leu Leu Met 410 1411 age aac tte tte gea cag ace aag gtt ttg get tte ggt aag aac get Ser Asn Phe Phe Ala Gln Thr Lys Val Leu Ala Phe Gly Lys Asn Ala 425 430

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440

1459

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Asn Arg Ala Glu Lys Tyr Thr Phe Ser Ala Ala Gly Leu His Val Asp	
35 40 45	
35 40 45 Leu Ser Lys Asn Leu Leu Asp Asp Ala Thr Leu Thr Lys Leu Leu Ala	
Leu Ser Lys Asn Leu Leu Asp Asp Ala Thr Leu Thr Lys Leu Leu Ala 50 55 60 Leu Thr Glu Glu Ser Gly Leu Arg Glu Arg Ile Asp Ala Met Phe Ala	
Leu Ser Lys Asn Leu Leu Asp Asp Ala Thr Leu Thr Lys Leu Leu Ala 50 Thr Glu Glu Ser Gly Leu Arg Glu Arg Ile Asp Ala Met Phe Ala 65 To 70 Thr Glu Asp Arg Ala Val Leu His Thr Ala	
Leu Ser Lys Asn Leu Leu Asp Asp Ala Thr Leu Thr Lys Leu Leu Ala  Leu Thr Glu Glu Ser Gly Leu Arg Glu Arg Ile Asp Ala Met Phe Ala 65 To Gly Glu His Leu Asn Asn Thr Glu Asp Arg Ala Val Leu His Thr Ala 90 Leu Arg Leu Pro Ala Glu Ala Asp Leu Ser Val Asp Gly Gln Asp Val	
Leu Ser Lys Asn Leu Leu Asp Asp Ala Thr Leu Thr Lys Leu Leu Ala 50 Thr Glu Glu Ser Gly Leu Arg Glu Arg Ile Asp Ala Met Phe Ala 65 Thr Glu His Leu Asn Asn Thr Glu Asp Arg Ala Val Leu His Thr Ala 95 Leu Arg Leu Pro Ala Glu Ala Asp Leu Ser Val Asp Gly Gln Asp Val 110 Asp Val Ala Ala Asp Val His Glu Val Leu Gly Arg Met Arg Asp Phe Ala Thr	

Ala Thr Lys Ala Leu Arg Ala Tyr Ala Thr Ala Gly Ile Ser Ala Glu Phe Val Ser Asn Val Asp Pro Ala Asp Leu Val Ser Val Leu Glu Asp Leu Asp Ala Glu Ser Thr Leu Phe Val Ile Ala Ser Lys Thr Phe Thr Thr Gln Glu Thr Leu Ser Asn Ala Arg Ala Arg Ala Trp Leu Val Glu Lys Leu Gly Glu Glu Ala Val Ala Lys His Phe Val Ala Val Ser 235 Thr Asn Ala Glu Lys Val Ala Glu Phe Gly Ile Asp Thr Asp Asn Met Phe Gly Phe Trp Asp Trp Val Gly Gly Arg Tyr Ser Val Asp Ser Ala 260 265 Val Gly Leu Ser Leu Met Ala Val Ile Gly Pro Arg Asp Phe Met Arg 280 Phe Leu Gly Gly Phe His Ala Met Asp Glu His Phe Arg Thr Thr Lys 295 Phe Glu Glu Asn Val Pro Ile Leu Met Ala Leu Leu Gly Val Trp Tyr Ser Asp Phe Tyr Gly Ala Glu Thr His Ala Val Leu Pro Tyr Ser Glu Asp Leu Ser Arg Phe Ala Ala Tyr Leu Gln Gln Leu Thr Met Glu Ser Asn Gly Lys Ser Val His Arg Asp Gly Ser Pro Val Ser Thr Gly Thr 355 Gly Glu Ile Tyr Trp Gly Glu Pro Gly Thr Asn Gly Gln His Ala Phe 375 Phe Gln Leu Ile His Gln Gly Thr Arg Leu Val Pro Ala Asp Phe Ile Gly Phe Ala Arg Pro Lys Gln Asp Leu Pro Ala Gly Glu Arg Thr Met His Asp Leu Leu Met Ser Asn Phe Phe Ala Gln Thr Lys Val Leu Ala Phe Gly Lys Asn Ala Glu Glu Ile Ala Ala Glu Gly Val Ala Pro Glu 440 Leu Val Asn His Lys Val Met Pro Gly Asn Arg Pro Thr Thr Thr Ile Leu Ala Glu Glu Leu Thr Pro Ser Ile Leu Gly Ala Leu Ile Ala Leu 470 475

Tyr Glu His Ile Val Met Val Gln Gly Val Ile Trp Asp Ile Asn Ser Phe Asp Gln Trp Gly Val Glu Leu Gly Lys Gln Gln Ala Asn Asp Leu Ala Pro Ala Val Ser Gly Glu Glu Asp Val Asp Ser Gly Asp Ser Ser Thr Asp Ser Leu Ile Lys Trp Tyr Arg Ala Asn Arg 535 <210> 43 <211> 630 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (1)..(630) <223> RXA01989 <400> 43 gtt aaa tca att cac aaa aca att cat gaa ggt act ggt gca ggt agt 48 Val Lys Ser Ile His Lys Thr Ile His Glu Gly Thr Gly Ala Gly Ser 5 15 1 10 gac ttc tta ggc tgg gtt gat tta cca gtt gat tac gac aaa gaa gaa 96 Asp Phe Leu Gly Trp Val Asp Leu Pro Val Asp Tyr Asp Lys Glu Glu 20 ttt tca aga att gtt gaa gca tca aaa cgc att aaa gaa aat tct gat Phe Ser Arg Ile Val Glu Ala Ser Lys Arg Ile Lys Glu Asn Ser Asp 35 gtt tta gta gtc atc ggt att ggt ggt tct tac tta ggt gca cgt gca 192 Val Leu Val Val Ile Gly Ile Gly Gly Ser Tyr Leu Gly Ala Arg Ala 50 qca atc gaa atg tta acg tca tca ttt aga aac agc aat gaa tac cct 240 Ala Ile Glu Met Leu Thr Ser Ser Phe Arg Asn Ser Asn Glu Tyr Pro 65 70 288 gaa att gta ttt gtt ggt aat cac tta tca tca aca tat acg aaa gag Glu Ile Val Phe Val Gly Asn His Leu Ser Ser Thr Tyr Thr Lys Glu 85 tta gtt gat tat tta gca gac aaa gat ttc tct gta aac gtt att tct 336 Leu Val Asp Tyr Leu Ala Asp Lys Asp Phe Ser Val Asn Val Ile Ser 100 384 aaa tot ggt aca act aca gaa cca gca gtt gca ttt aga ttg ttc aaa Lys Ser Gly Thr Thr Thr Glu Pro Ala Val Ala Phe Arg Leu Phe Lys 115 120 caa tta gtt gaa gaa aga tac ggt aaa gaa gaa gca caa aaa cgt ata 432 Gln Leu Val Glu Glu Arg Tyr Gly Lys Glu Glu Ala Gln Lys Arg Ile 130 ttt gca aca acg gat aaa gaa aaa ggt gct tta aaa cag ttg gct aca 480

Phe Ala Thr 145	Thr Asp	Lys Glu 150	Lys (	Gly F		Leu 155	Lys	Gln	Leu	Ala	Thr 160	
aac gaa ggt Asn Glu Gly		Thr Phe		Val E								528
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tta tct Leu Ser 210												630
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Phe Ser Arg	Ile Val	Glu Ala	Ser :	Lys A	Arg ]	Ile	Lys	Glu 45	Asn	Ser	Asp	
Val Leu Val 50	Val Ile	Gly Ile	-	Gly S	Ser I	Tyr	Leu 60	Gly	Ala	Arg	Ala	
Ala Ile Glu 65	Met Leu	Thr Ser	Ser :	Phe F	Arg A	Asn 75	Ser	Asn	Glu	Tyr	Pro 80	
Glu Ile Val	Phe Val 85	_	His :				Thr	Tyr	Thr	Lys 95	Glu	
Leu Val Asp	Tyr Leu 100	Ala Asp	_	Asp F 105	Phe S	Ser	Val	Asn	Val 110	Ile	Ser	
Lys Ser Gly 115	Thr Thr	Thr Glu	Pro 1	Ala V	Val <i>P</i>	Ala	Phe	Arg 125	Leu	Phe	Lys	
Gln Leu Val 130	Glu Glu	Arg Tyr		Lys 0	Glu (		Ala 140	Gln	Lys	Arg	Ile	
Phe Ala Thr 145	Thr Asp	Lys Glu 150	Lys (	Gly A		Leu 155	Lys	Gln	Leu	Ala	Thr 160	
Asn Glu Gly	Tyr Glu 165		Ile '		Pro <i>P</i> 170	Asp .	Asp	Val	Gly	Gly 175	Arg	
Tyr Ser Val	Leu Thr 180	Ala Val		Leu I 185	Leu F	Pro	Ile	Ala	Thr 190	Ala	Gly	

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Leu Ser 210						
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gag gcc gac gg Glu Ala Asp Gl						
acc gtg gtg ta Thr Val Val Ty 2	r Ser Asp A					
tat gtt gtc gg Tyr Val Val Gl 40						
cta ctc gaa gg Leu Leu Glu Gl 55						
gat gtc tac at Asp Val Tyr Me 70	t Asp Ser I		Val Glu	Gln Met		
tgg aaa atc aa Trp Lys Ile Ly						
gag att gct tc Glu Ile Ala Se 10	r Glu Ile G					
gag aaa aac aa Glu Lys Asn Ly 120						
gca gcg gca gg Ala Ala Ala Gl 135	y Lys Pro V					

		tca gag aaa gaa gac ctc a Ser Glu Lys Glu Asp Leu A 160	
	Thr Asn Trp	aac ggc gca acc aca gat o Asn Gly Ala Thr Thr Asp I 175	
cgt ttc ttg ttg ct Arg Phe Leu Leu Leu 185	cgc cac ggc 1 Arg His Gly	caa act gct atg tca gtg g Gln Thr Ala Met Ser Val 1 190 195	gca cgc 691 Ala Arg
ctt tac tcc ggt age Leu Tyr Ser Gly Are 200	g tcc aac cca g Ser Asn Pro 205	gag ctg tct gaa ctt ggt g Glu Leu Ser Glu Leu Gly G 210	gaa aaa 739 Glu Lys
caa gca gca gcg gc Gln Ala Ala Ala Al 215	a gca cga cga a Ala Arg Arg 220	ctc gct caa acc ggt ggc a Leu Ala Gln Thr Gly Gly 3 225	atc gac 787 Ile Asp
		cgc acg atg caa acc gca o Arg Thr Met Gln Thr Ala o 240	
gca gcg gcc gca ct Ala Ala Ala Ala Le 25	u Gly Met Lys	gta cgt gtt atc gat gat c Val Arg Val Ile Asp Asp I 255	ctc atc 883 Leu Ile 260
		gga aaa tca ttt tca gaa g Gly Lys Ser Phe Ser Glu A 270 275	
		aag tgg ctc act gac tca t Lys Trp Leu Thr Asp Ser S 290	
		cag acg gtt aat cga cgt of Gln Thr Val Asn Arg Arg N 305	
		gaa tac ggt gca gcg aat g Glu Tyr Gly Ala Ala Asn V 320	
	l Thr Pro Ile	aaa gcc atc atg agg caa g Lys Ala Ile Met Arg Gln A 335	
gac gca ggc cca tc Asp Ala Gly Pro Se 345	c ttc ttt cag r Phe Phe Gln	aag gca cac ctt gac ttg c Lys Ala His Leu Asp Leu A 350 355	gcg tcg 1171 Ala Ser
		gac ggc cca acc tgc gta a Asp Gly Pro Thr Cys Val A 370	
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ctc			1269

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Leu Lys Glu Ile Ala Tyr Val Val Gly Thr Lys Ala Thr Asn Asn Val 35 40 45

Ala Glu Tyr Arg Gly Leu Leu Glu Gly Leu Lys Ala Arg Glu Leu 50 55 60

Gly Ala Thr Ser Val Asp Val Tyr Met Asp Ser Lys Leu Val Val Glu 65 70 75 80

Gln Met Ser Gly Arg Trp Lys Ile Lys His Pro Asp Met Lys Val Leu 85 90 95

Ala Ile Glu Ala Lys Glu Ile Ala Ser Glu Ile Gly Ser Val Ser Tyr 100 105 110

Thr Trp Ile Pro Arg Glu Lys Asn Lys Arg Ala Asp Ala Leu Ser Asn 115 120 125

Val Ala Met Asp Ala Ala Ala Gly Lys Pro Val Gly Val Val Gly 130 135 140

Asp Ser Ala Ser Val Ser Ser Ala Ser Ser Val Ala Gly Ser Glu Lys 145 150 155 160

Glu Asp Leu Asn Cys Thr Glu Thr Lys Pro Thr Asn Trp Asn Gly Ala 165 170 175

Thr Thr Asp Pro Thr Arg Phe Leu Leu Arg His Gly Gln Thr Ala 180 185 190

Met Ser Val Ala Arg Leu Tyr Ser Gly Arg Ser Asn Pro Glu Leu Ser 195 200 205

Glu Leu Gly Glu Lys Gln Ala Ala Ala Ala Ala Arg Arg Leu Ala Gln 210 215 220

Thr Gly Gly Ile Asp Ala Ile Val Ser Ser Pro Leu Thr Arg Thr Met 225 230 235 240

Gln Thr Ala Glu Ala Ala Ala Ala Leu Gly Met Lys Val Arg Val 245 250 255

Ile Asp Asp Leu Ile Glu Thr Asp Phe Gly Leu Trp Asp Gly Lys Ser 260 265 270

Phe Ser Glu Ala His Glu Gln Asp Pro Glu Leu His Thr Lys Trp Leu 275 280 285

Thr Asp Ser 290	Ser Val	Ala Pro 295	Pro G	ly Gly		Ser Leu 300	Gln	Thr	Val					
Asn Arg Arg 305		Lys Ala 310	Arg G	lu Ser	Leu ( 315	Gln Arg	Glu	Tyr	Gly 320					
Ala Ala Asn	Val Leu 325	Val Val	Ser Hi	is Val 330	Thr I	Pro Ile	Lys	Ala 335	Ile					
Met Arg Gln	Ala Leu 340	Asp Ala		ro Ser 45	Phe I	Phe Gln	Lys 350	Ala	His					
Leu Asp Leu 355	Ala Ser	Leu Ser	Ile Al	la Glu	Phe 7	Tyr Glu 365	Asp	Gly	Pro					
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					acct a		aac	gga	aaa	60 115				
gctgtacaac	cgtggaaca	t aaagt	ggcaa a	actagta gc gaa	icct a	atg act Met Thr 1 aac gca	aac Asn	gga Gly aac	aaa Lys 5 cag					
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gctgtacaac aaacgcgcgt ttg att ctt Leu Ile Leu ttc act gga	cgtggaaca ctt cgt Leu Arg 10 tgg gtc Trp Val 25 gtc ctc	cac ggt His Gly gac gtc Asp Val	cag aq Gln Se aat ct Asn Le	gc gaa er Glu 15 tg acc eu Thr 30	tgg a Trp 1 gaa ( Glu (	atg act Met Thr 1 aac gca Asn Ala cag ggt Gln Gly	aac Asn tcc Ser gag Glu 35	gga Gly aac Asn 20 gct Ala	aaa Lys 5 cag Gln gag Glu	115 163				
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gcg gat gac gca gag tac tcc cag gca aat gac cct cgt tac gcg gac Ala Asp Asp Ala Glu Tyr Ser Gln Ala Asn Asp Pro Arg Tyr Ala Asp 120 125 130	499
ctc gac gta gtt cca cgc acc gaa tgc ctc aag gac gtt gtg gtt cgt Leu Asp Val Val Pro Arg Thr Glu Cys Leu Lys Asp Val Val Val Arg 135 140 145	547
ttt gtt cct tac ttc gag gaa gaa atc ctg cca cgc gca aag aag ggc Phe Val Pro Tyr Phe Glu Glu Glu Ile Leu Pro Arg Ala Lys Lys Gly 150 155 160 165	595
gaa acc gtc ctc atc gca gca cac ggc aac tcc ctg cgt gcg ctg gtt Glu Thr Val Leu Ile Ala Ala His Gly Asn Ser Leu Arg Ala Leu Val 170 175 180	643
aag cac ctt gac ggc atc tcc gat gct gat atc gca gag ctc aac atc Lys His Leu Asp Gly Ile Ser Asp Ala Asp Ile Ala Glu Leu Asn Ile 185 190 195	691
cca acc ggc atc cca ctg gtc tac gaa atc gcc gaa gac ggt tcc gta Pro Thr Gly Ile Pro Leu Val Tyr Glu Ile Ala Glu Asp Gly Ser Val 200 205 210	739
gta aac cca ggc ggc acc tac ctc gat cct gag gca gca gcc ggc Val Asn Pro Gly Gly Thr Tyr Leu Asp Pro Glu Ala Ala Ala Ala Gly 215 220 225	787
gca gca gca gta gca aac cag ggt aat aag tagctatttg taggtgagca Ala Ala Ala Val Ala Asn Gln Gly Asn Lys	837
230 235	
	840
230 235	840
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				85					90					95		
Lys	Tyr	Gly	Asp 100	Asp	Gln	Phe	Met	Glu 105	Trp	Arg	Arg	Ser	Tyr 110	Asp	Thr	
Pro	Pro	Pro 115	Glu	Leu	Ala	Asp	Asp 120	Ala	Glu	Tyr	Ser	Gln 125	Ala	Asn	Asp	
Pro	Arg 130	Tyr	Ala	Asp	Leu	Asp 135	Val	Val	Pro	Arg	Thr 140	Glu	Cys	Leu	Lys	
Asp 145	Val	Val	Val	Arg	Phe 150	Val	Pro	Tyr	Phe	Glu 155	Glu	Glu	Ile	Leu	Pro 160	
Arg	Ala	Lys	Lys	Gly 165	Glu	Thr	Val	Leu	Ile 170	Ala	Ala	His	Gly	Asn 175	Ser	
Leu	Arg	Ala	Leu 180	Val	Lys	His	Leu	Asp 185	Gly	Ile	Ser	Asp	Ala 190	Asp	Ile	
Ala	Glu	Leu 195	Asn	Ile	Pro	Thr	Gly 200	Ile	Pro	Leu	Val	Tyr 205	Glu	Ile	Ala	
Glu	Asp 210	Gly	Ser	Val	Val	Asn 215	Pro	Gly	Gly	Thr	Tyr 220	Leu	Asp	Pro	Glu	
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														atc Ile 20		163
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														tac Tyr		259
gcg Ala	gcc Ala	tcc Ser	cca Pro	ttg Leu	cag Gln	cgt Arg	gtg Val	cag Gln	gaa Glu	acc Thr	tcc Ser	gaa Glu	ccg Pro	ttc Phe	atc Ile	307

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Asp Val Thr Tyr Ile Ala Ala Ser Pro Leu Gln Arg Val Gln Glu Thr

Ser Glu Pro Phe Ile Lys Val Thr Gly Leu Glu Leu Ile Thr Asp Glu

Asp Leu Leu Glu Ala Gly Asn Arg Phe Glu Gly Leu Arg Thr Lys Gly

Trp	Arg	Ser	Gln 100	Leu	Trp	Asn	Pro	Val 105	Arg	Trp	Pro	Leu	Met 110	Tyr	Asn	
Pro	Thr	Leu 115	Pro	Ser	Trp	Gly	Glu 120	His	Tyr	Thr	Asp	Ile 125	Leu	Glu	Arg	
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Ala	Arg	Gly	Gln	Ser 165	Leu	Ser	His	Asn	Pro 170	Ala	Thr	Arg	Gln	Cys 175	Asp	
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ccti	tttt	ta a	agtg	ggcgg	gt ca	aggaa	attt	te	gcaca	aggt				gtç Val		115
					gca Ala											163
					ggg Gly											211
					tta Leu											259
					cgt Arg											307
					cgt Arg 75											355

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•	-		gcg tca att tcc Ala Ser Ile Ser 195	
			cc cat atg ctg Ser His Met Leu 210	-
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Gly Gln Val Leu 35	Pro Gly Gln	Thr Pro Gly I	Leu His Leu Ser 7 45	Asp Lys
Gly Glu Glu Gln 50	Ala Arg Glu 55	Val Ala Gln A	Arg Leu Ala Glu ' 60	Val Pro
Ile Thr Ala Val 65	Tyr Ser Ser 70	Pro Met Glu A	Arg Ala Gln Glu' 75	Thr Ala 80

Ala Pro Thr Val Ser Ala His Gly Leu Glu Leu Thr Val Glu Pro Gly 90 Leu Ile Glu Cys Asp Phe Gly Glu Trp Thr Gly Arg Lys Leu Thr Glu 105 Leu Asn Ala Leu Glu Glu Trp Lys Ala Val Gln Lys Thr Pro Ser Thr 120 Phe Arg Phe Pro Gly Gly Glu Ser Phe Val Glu Met Gln Asp Arg Met 135 Val Glu Ala Ile Gly Asn Ile Ala Gln Gln His Pro Gly Glu Ile Val 150 155 Ala Ala Phe Ser His Ala Asp Thr Ile Lys Ala Ala Val Ala His Phe 170 Val Gly Thr Pro Leu Asp Ser Phe Gln Arg Ile Phe Ile Asp Thr Ala 180 185 Ser Ile Ser Ala Val Glu Phe Thr Gly Lys Ser Ser Gly Val Ser Ser 200 His Met Leu Leu Thr Asn Ser Arg Thr Gly Ser Leu Gly Tyr Leu Arg 215 Asp Lys Leu Pro Lys Ala Pro Gln Pro 230 <210> 53 <211> 1161 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(1138) <223> RXA00206 <400> 53 ttaaataaga tggtcagaga cagttttttg gcctgtcaac ccctgtgatt ctcttatttt 60 tgggtgattg ttccggcgcg ggtgttgtga tgggtttaat atg gaa gac atg cga Met Glu Asp Met Arg att gct act ctc acg tca ggc ggc gac tgc ccc gga cta aac gcc gtc 163 Ile Ala Thr Leu Thr Ser Gly Gly Asp Cys Pro Gly Leu Asn Ala Val 211 ate ega gga ate gte ege aca gee age aat gaa tit gge tee ace gte Ile Arg Gly Ile Val Arg Thr Ala Ser Asn Glu Phe Gly Ser Thr Val gtt ggt tat caa gac ggt tgg gaa gga ctg tta ggc gat cgt cgc gta 259 Val Gly Tyr Gln Asp Gly Trp Glu Gly Leu Leu Gly Asp Arg Arg Val 45

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						gga Gly										451
						gtc Val										499
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						cgc Arg										595
						gtc Val										643
						ggc Gly										69Ì
-			-		-	gag Glu		_	_		_	-	-	-		739
			Ğlu	Lys	Tyr	ggc Gly 220	Ile	Ile	Val	Val	Ala	Ğlu				787
						gag Glu										835
						gga Gly										883
						gat Asp										931
						act Thr										979
tat	ggt	gtt	cgt	gca	gct	cgt	gcg	tgc	cat	gag	gga	agc	ttt	gac	aag	1027

Tyr Gly Val Arg Ala Ala Arg Ala Cys His Glu Gly Ser Phe Asp Lys gtt gtt gct ttg aag ggt gag agc att gag atg atc acc ttt gaa gaa 1075 Val Val Ala Leu Lys Gly Glu Ser Ile Glu Met Ile Thr Phe Glu Glu qca qtc qqa acc ttq aaq qaa gtt cca ttc gaa cgc tgg gtt act gcc Ala Val Gly Thr Leu Lys Glu Val Pro Phe Glu Arg Trp Val Thr Ala 335 cag gca atg ttt gga tagtttttcg ggcttttatc aac 1161 Gln Ala Met Phe Gly 345 <210> 54 <211> 346 <212> PRT <213> Corynebacterium glutamicum <400> 54 Met Glu Asp Met Arg Ile Ala Thr Leu Thr Ser Gly Gly Asp Cys Pro Gly Leu Asn Ala Val Ile Arg Gly Ile Val Arg Thr Ala Ser Asn Glu Phe Gly Ser Thr Val Val Gly Tyr Gln Asp Gly Trp Glu Gly Leu Leu Gly Asp Arg Arg Val Gln Leu Tyr Asp Asp Glu Asp Ile Asp Arg Ile 55 Leu Leu Arg Gly Gly Thr Ile Leu Gly Thr Gly Arg Leu His Pro Asp Lys Phe Lys Ala Gly Ile Asp Gln Ile Lys Ala Asn Leu Glu Asp Ala Gly Ile Asp Ala Leu Ile Pro Ile Gly Gly Glu Gly Thr Leu Lys Gly Ala Lys Trp Leu Ser Asp Asn Gly Ile Pro Val Val Gly Val Pro Lys 115 Thr Ile Asp Asn Asp Val Asn Gly Thr Asp Phe Thr Phe Gly Phe Asp Thr Ala Val Ala Val Ala Thr Asp Ala Val Asp Arg Leu His Thr Thr Ala Glu Ser His Asn Arg Val Met Ile Val Glu Val Met Gly Arg His 170 Val Gly Trp Ile Ala Leu His Ala Gly Met Ala Gly Gly Ala His Tyr Thr Val Ile Pro Glu Val Pro Phe Asp Ile Ala Glu Ile Cys Lys Ala

205

200

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Ala 225	Glu	Gly	Ala	Leu	Pro 230	Arg	Glu	Gly	Thr	Met 235	Glu	Leu	Arg	Glu	Gly 240	
His	Ile	Asp	Gln	Phe 245	Gly	His	Lys	Thr	Phe 250	Thr	Gly	Ile	Gly	Gln 255	Gln	
Ile	Ala	Asp	Glu 260	Ile	His	Val	Arg	Leu 265	Gly	His	Asp	Val	Arg 270	Thr	Thr	
Val	Leu	Gly 275	His	Ile	Gln	Arg	Gly 280	Gly	Thr	Pro	Thr	Ala 285	Phe	Asp	Arg	
Val	Leu 290	Ala	Thr	Arg	Tyr	Gly 295	Val	Arg	Ala	Ala	Arg 300	Ala	Cys	His	Glu	
Gly 305	Ser	Phe	Asp	Lys	Val 310	Val	Ala	Leu	Lys	Gly 315	Glu	Ser	Ile	Glu	Met 320	
Ile	Thr	Phe	Glu	Glu 325	Ala	Val	Gly	Thr	Leu 330	Lys	Glu	Val	Pro	Phe 335	Glu	
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	att Ile															211
	ttt Phe															259
		10														

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														gcc Ala		691
														gat Asp		739
tcc Ser	ccg Pro 215	act Thr	gtg Val	gcg Ala	gca Ala	gcg Ala 220	cgc Arg	gct Ala	tta Leu	att Ile	gat Asp 225	agc Ser	ggt Gly	gtc Val	acc Thr	787
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														cag Gln 260		883
														tcc Ser		931
														tac Tyr		979
														gat Asp		1027
ctt	cgg	gcg	gag	cac	gtg	gtc	atc	aaa	tcg	ctt	taga	acca	ege a	aaaa	agcctc	1080

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aaa 1083

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<211> 320

<212> PRT

<213> Corynebacterium glutamicum

<400> 56

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Ser Thr Val Ala Gly Gly Phe Gly Thr Gly Val Ala Ala Thr Leu Phe 35 40 45

Tyr Gly Gly Asn Glu Thr Phe Ala Val Phe Pro Ala Pro Glu Ile Ser 50 55 60

His Tyr Met Arg Leu Val Thr Phe Ala Gly Leu Pro His Glu Ile Ile 65 70 75 80

Pro Val Ala Gly Pro Ile Pro Met His Leu Thr Met Arg Asp Ala Glu 85 90 95

Gly Asn Glu Thr Lys Phe Lys Asp Ser Pro Met Pro Leu Asp Val Ser 100 105 110

Gln Leu Ala Ile Leu Arg Asp Leu Val Val Arg Arg Ala Glu Asp Ala 115 120 125

Ala Trp Val Leu Leu Gly Gly Asn Leu Pro Ser Ile Ala Pro Ala Ala 130 135 140

Trp Phe Val Asp Val Val Arg Ser Leu Arg Leu Tyr His Pro His Val 145 150 155 160

Lys Val Ala Ile Ala Ala Thr Gly Ala Ala Leu Arg Ala Val Ile Arg 165 170 175

Gln Leu Ala Ala Thr Ser Pro Asp Ala Leu Ile Val Ala Ala Glu Glu 180 185 190

Ile Glu Ile Ala Thr Gly Leu Glu Pro Lys Thr Leu Arg Gly Pro Trp 195 200 205

Val Glu Gly Asp Leu Ser Pro Thr Val Ala Ala Ala Arg Ala Leu Ile 210 215 220

Asp Ser Gly Val Thr Glu Val Leu Val Thr Asn Lys Arg Thr Glu Ser 225 230 235 240

Leu Tyr Val Ser Glu Ser Glu Ser Leu Leu Ala Ser Tyr Asp Ser Thr 245 250 255

Pro Gly Lys Gln Gly Val Asn Trp Arg Glu Ser Phe Thr Ala Gly Phe

270

Leu Ala Ala Ser Asn Asp Gly Lys Ser Thr Glu Asp Ser Val Ile Asn 275

Ala Val Ala Tyr Ala Asn Ala Glu Gly Ser Glu Trp Asp Asn Tyr Ile

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gaa aag gtg ctt atc gac gcg ctc cgc ccc gaa gtc acc tgg gtt gtc

499

Glu	Lys	Val 120	Leu	Ile	Asp	Ala	Leu 125	Arg	Pro	Glu	Val	Thr 130	Trp	Val	Val	
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	acc Thr															595
gat Asp	acc Thr	tca Ser	gac Asp	aag Lys 170	cca Pro	ctg Leu	atg Met	gcg Ala	ttg Leu 175	ggc Gly	gag Glu	agc Ser	ttg Leu	gat Asp 180	aca Thr	643
	ggc Gly															691
	ctg Leu															739
ggc Gly	gat Asp 215	tac Tyr	gac Asp	gcc Ala	atc Ile	atc Ile 220	gca Ala	gct Ala	gcg Ala	gac Asp	gta Val 225	ctg Leu	gtt Val	aac Asn	cgt Arg	787
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	aac Asn															883
	tcc Ser															931
	cgt Arg															979
	tac Tyr 295															1027
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310

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acc Thr	tcc Ser	cca Pro	gaa Glu 185	gac Asp	ttt Phe	gag Glu	aag Lys	acc Thr 190	atc Ile	gat Asp	gca Ala	atc Ile	ggc Gly 195	acc Thr	ggt Gly	691
gag Glu	aag Lys	ggc Gly 200	cgc Arg	tac Tyr	ctg Leu	cta Leu	gca Ala 205	gct Ala	acc Thr	ttc Phe	ggt Gly	aac Asn 210	gtc Val	cac His	ggc Gly	739
					aac Asn											787
					cgc Arg 235											835
					ttc Phe											883
					acc Thr											931
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aac Asn	tac Tyr 295	aac Asn	ggc	gtt Val	ctc Leu	aag Lys 300	atc Ile	gac Asp	ggc Gly	gag Glu	gtc Val 305	gga Gly	aac Asn	aag Lys	aag Lys	1027
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Ile	Ile 50	Gln	Phe	Ser	Thr	Gly 55	Gly	Ala	Glu	Phe	Gly 60	Ser	Gly	Leu	Ala
Val 65	Lys	Asn	Lys	Val	Lys 70	Gly	Ala	Val	Ala	Leu 75	Ala	Ala	Phe	Ala	His 80
Glu	Ala	Ala	Lys	Ser 85	Tyr	Gly	Ile	Asn	Val 90	Ala	Leu	His	Thr	Asp 95	His
Cys	Gln	Lys	Glu 100	Val	Leu	Asp	Glu	Tyr 105	Val	Arg	Pro	Leu	Leu 110	Ala	Ile
Ser	Gln	Glu 115	Arg	Val	Asp	Arg	Gly 120	Glu	Leu	Pro	Leu	Phe 125	Gln	Ser	His
Met	Trp 130	Asp	Gly	Ser	Ala	Val 135	Pro	Ile	Asp	Glu	Asn 140	Leu	Glu	Ile	Ala
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Glu 225	Val	Leu	Leu	Glu	Gly 230	Gln	Gln	Val	Ala	Arg 235	Lys	Lys	Leu	Gly	Let 240
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Val 305	Gly	Asn	Lys	Lys	Ala 310	Tyr	Asp	Pro	Arg	Ser 315	Tyr	Met	Lys	Lys	Ala 320
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		ggt Gly														787
_		gac Asp	_	_				_			-			-		835
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Lys	Glu	Tyr 35	Phe	Glu	Lys	Val	Asp 40	Val	Ala	Val	Thr	Val 45	Pro	Phe	Thr	
Asp	Ile 50	Arg	Ser	Val	Gln	Thr 55	Leu	Val	Glu	Gly	Asp 60	Lys	Leu	Glu	Val	
Thr 65	Phe	Gly	Ala	Gln	Asp 70	Val	Ser	Gln	His	Glu 75	Ser	Gly	Ala	Tyr	Thr 80	
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Val	Val	Gly	His 100	Ser	Glu	Arg	Arg	Glu 105	Tyr	His	Asn	Glu	Ser 110	Asp	Glu	
Leu	Val	Ala 115	Ala	Lys	Ala	Lys	Ala 120	Ala	Leu	Ser	Asn	Gly 125	Ile	Ser	Pro	
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Val 145	Glu	Tyr	Val	Val	Glu 150	Gln	Thr	Arg	Lys	Ser 155	Leu	Ala	Gly	Leu	Asp 160	
Ala	Ala	Glu	Leu	Ala 165	Asn	Thr	Val	Ile	Ala 170	Tyr	Glu	Pro	Val	Trp 175	Ala	

Ile Gly Thr Gly Lys Val Ala Ser Ala Ala Asp Ala Gln Glu Val Cys Lys Ala Ile Arg Gly Leu Ile Val Glu Leu Ala Gly Asp Glu Val Ala 200 Glu Gly Leu Arg Ile Leu Tyr Gly Gly Ser Val Lys Ala Glu Thr Val Ala Glu Ile Val Gly Gln Pro Asp Val Asp Gly Gly Leu Val Gly Gly Ala Ser Leu Asp Gly Glu Ala Phe Ala Lys Leu Ala Ala Asn Ala Ala 250 Ser Val Ala <210> 63 <211> 1563 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(1540) <223> RXN01225 <400> 63 tttgggctaa tgttggggg agtgctttca actatccacg agagctgccc agtgataaac 60 cccgggttaa ccccacgcct aagtcagtga aggacttttt atg acg cac aac cac 115 Met Thr His Asn His aag gac tgg aac gat cgc att gca gtt gcg gag gaa atg gtg ccg ttg 163 Lys Asp Trp Asn Asp Arg Ile Ala Val Ala Glu Glu Met Val Pro Leu 15 atc ggg cgc ctg cac cgc aac aac aac gtg gtg gtt tcc gta ttc ggt 211 Ile Gly Arg Leu His Arg Asn Asn Val Val Val Ser Val Phe Gly cgt ctc ctt gtg aat gtc tca gac atc gat atc atc aag tct cac cgc 259 Arg Leu Leu Val Asn Val Ser Asp Ile Asp Ile Ile Lys Ser His Arg 45 307 tac gcc cgc cac atc ata tcc aag gaa ctt cca ctg gaa agc tcc ttg Tyr Ala Arg His Ile Ile Ser Lys Glu Leu Pro Leu Glu Ser Ser Leu gat att ttg cgc gaa ctg gta gat atg aac ctt ggt acc gca tcg atc 355 Asp Ile Leu Arg Glu Leu Val Asp Met Asn Leu Gly Thr Ala Ser Ile 403 gac ctq qqa caq ctg qcc tac aqc ttc qaa qaa tcc qaa agc acc gac Asp Leu Gly Gln Leu Ala Tyr Ser Phe Glu Glu Ser Glu Ser Thr Asp 90 95

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	-	-	-	-	gta Val				_		_		-	-	-	883
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-				-	cgc Arg				_				_	-		1027
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Thr Gly Ala	Ala Lys 345	Ala	Val	Ser	Lys 350	Ala	Leu	Pro	Glu	Leu 355	Glu	Gly	
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gct gtg ctc Ala Val Leu 375	-		_		-			-	-	-		-	1267
aac gag ttc Asn Glu Phe 390	-	_			-			-	_	-	-		1315
atc gac tgg Ile Asp Trp		Ser											1363
acc acc cac Thr Thr His													1411
cgc cac ctg Arg His Leu 440													1459
aac cag gtc	_		-				-			_		-	1507
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Asn Gln Val 455 gtg tac ccg Val Tyr Pro	gag cgc	agg Arg	460 cag	cca	gcc	gta	cta Leu	465					1560 1563
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Gly Thr Ala Ser Ile Asp Leu Gly Gln Leu Ala Tyr Ser Phe Glu Glu 85 90 95

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Gly	Phe 130	Gly	Arg	Ile	Gly	Arg 135	Leu	Leu	Ala	Arg	Ile 140	Leu	Val	Ser	Arg
Glu 145	Ala	Leu	Tyr	Asp	Gly 150	Ala	Arg	Leu	Arg	Ala 155	Ile	Val	Val	Arg	Lys 160
Asn	Gly	Glu	Glu	Asp 165	Leu	Val	Lys	Arg	Ala 170	Ser	Leu	Leu	Arg	Arg 175	Asp
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Asp	Pro 210	Ala	Thr	Ile	Asp	Tyr 215	Thr	Glu	Tyr	Gly	Ile 220	Asn	Asp	Ala	Val
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Pro	Asp 370	Val	Ser	Met	Ala	Val 375	Leu	Asn	Leu	Thr	Leu 380	Asn	Thr	Glu	Val
Asp 385	Arg	Asp	Glu	Val	Asn 390	Glu	Phe	Leu	Arg	Arg 395	Val	Ser	Leu	His	Ser 400
Asp	Leu	Arg	Gln	Gln 405	Ile	Asp	Trp	Ile	Arg 410	Ser	Pro	Glu	Val	Val 415	Ser

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Thr Ile Ala Thr Gly Arg His Leu Val Leu Tyr Val Trp Tyr Asp Asn 435

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92

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ggt Gly	cgc Arg 135	ctg Leu	ctg Leu	gcc Ala	cgc Arg	atc Ile 140	ctg Leu	gtt Val	tcc Ser	cgc Arg	gag Glu 145	gca Ala	ctg Leu	tat Tyr	gac Asp	547
ggt Gly 150	gct Ala	cgt Arg	ctg Leu	cgc Arg	gcc Ala 155	atc Ile	gtg Val	gtc Val	cgc Arg	aaa Lys 160	aat Asn	ggt Gly	gaa Glu	gaa Glu	gac Asp 165	595
	gtc Val															643
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	ggc Gly															739
	tac Tyr 215															787
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	gtt Val															883
	atc Ile															931
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	cac His															1075
	tct Ser															1123
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ac 1563	•
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2210> 66 (2211> 480) (2212> PRT) (2213> Corynebacterium glutamicum) (400> 66 (et Thr His Asn His Lys Asp Trp Asn Asp Arg Ile Ala Val Ala Glu 1 5 10 15 (ilu Met Val Pro Leu Ile Gly Arg Leu His Arg Asn Asn Asn Val Val 20 25 30 (val Ser Val Phe Gly Arg Leu Leu Val Asn Val Ser Asp Ile Asp Ile	
2210> 66 2211> 480 2212> PRT 2213> Corynebacterium glutamicum 2400> 66 Met Thr His Asn His Lys Asp Trp Asn Asp Arg Ile Ala Val Ala Glu 1 5 10 15 Slu Met Val Pro Leu Ile Gly Arg Leu His Arg Asn Asn Asn Val Val 20 25 30  Val Ser Val Phe Gly Arg Leu Leu Val Asn Val Ser Asp Ile Asp Ile 35 40 45  Sle Lys Ser His Arg Tyr Ala Arg His Ile Ile Ser Lys Glu Leu Pro	
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Glu 145	Ala	Leu	Tyr	Asp	Gly 150	Ala	Arg	Leu	Arg	Ala 155	Ile	Val	Val	Arg	Lys 160
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Ser	Val	His	Gly 180	Gly	Phe	Asp	Gly	Thr 185	Ile	Thr	Thr	Asp	Tyr 190	Asp	Asn
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Thr	Ala	Asp 275	Asp	Gln	Ile	Val	Ser 280	Ala	Ala	Ser	Cys	Thr 285	Thr	Asn	Ala
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tac act ggc gac Tyr Thr Gly Asp 185	cag cgc ctg Gln Arg Leu	cac gat gca His Asp Ala 190	Pro His Arg A	ac ctg cgt sp Leu Arg 95	691
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gaa gag cca ctg Glu Glu Pro Leu 280	gtt tcc acc Val Ser Thr	gac atc gtc Asp Ile Val 285	cac gat tcc c His Asp Ser H 290	ac ggc tcc is Gly Ser	979
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	gcc tcc cta cca acc c Ala Ser Leu Pro Thr L 45	
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	g tac gtt gca ctt gcc g n Tyr Val Ala Leu Ala A ) 95	
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	c ttc gac cca cgc gaa a g Phe Asp Pro Arg Glu T 125	
	c gct cag gag ctc gca g e Ala Gln Glu Leu Ala A 140	
	t gac ggc ttc ggt gtt g r Asp Gly Phe Gly Val V 155	
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ggt gtt att ga Gly Val Ile Gl 215		a Ala Lys Ala			787
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atc cca gaa gg Ile Pro Glu Gl 295		Leu Asp Ile			1027
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<213> Corynebacterium glutamicum

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Met Ala Val Lys Thr Leu Lys Asp Leu Leu Asp Glu Gly Val Asp Gly 1 5 10 15

Arg His Val Ile Val Arg Ser Asp Phe Asn Val Pro Leu Asn Asp Asp 20 25 30

Arg Glu Ile Thr Asp Lys Gly Arg Ile Ile Ala Ser Leu Pro Thr Leu  $35 \hspace{1.5cm} 40 \hspace{1.5cm} 45$ 

Lys Ala Leu Ser Glu Gly Gly Ala Lys Val Ile Val Met Ala His Leu 50 60

Gly Arg Pro Lys Gly Glu Val Asn Glu Lys Tyr Ser Leu Ala Pro Val 65 70 75 80

Ala Glu Ala Leu Ser Asp Glu Leu Gly Gln Tyr Val Ala Leu Ala Ala 85 90 95

Asp Val Val Gly Glu Asp Ala His Glu Arg Ala Asn Gly Leu Thr Glu 100 105 110

Gly Asp Ile Leu Leu Glu Asn Val Arg Phe Asp Pro Arg Glu Thr 115 120 125

Ser Lys Asp Glu Ala Glu Arg Thr Ala Phe Ala Gln Glu Leu Ala Ala 130 135 140

Leu Ala Ala Asp Asn Gly Ala Phe Val Ser Asp Gly Phe Gly Val Val 145 150 155 160

His Arg Ala Gln Thr Ser Val Tyr Asp Ile Ala Lys Leu Leu Pro His 165 170 175

Tyr Ala Gly Gly Leu Val Glu Thr Glu Ile Ser Val Leu Glu Lys Ile 180 185 190

Ala Glu Ser Pro Glu Ala Pro Tyr Val Val Leu Gly Gly Ser Lys 195 200 205

Val Ser Asp Lys Ile Gly Val Ile Glu Ala Leu Ala Ala Lys Ala Asp 210 215 220

Lys Ile Ile Val Gly Gly Gly Met Cys Tyr Thr Phe Leu Ala Ala Gln 225 230 235 240

Gly His Asn Val Gln Gln Ser Leu Leu Gln Glu Glu Met Lys Ala Thr 245 250 255

Cys Thr Asp Leu Leu Ala Arg Phe Gly Asp Lys Ile Val Leu Pro Val 260 265 270

Asp Leu Val Ala Ala Ser Glu Phe Asn Lys Asp Ala Glu Lys Gln Ile 275 280 285

290	Ser Ile	Pro Glu 295	Gly Trp	Met Se 30		Asp	Ile	Gly	
Pro Glu Ser Val 305	Lys Asn 310	Phe Gly	Glu Val	Leu Se 315	r Thr	Ala	Lys	Thr 320	
Ile Phe Trp Asn	Gly Pro 325	Met Gly	Val Phe 330	Glu Ph	e Ala	Ala	Phe 335	Ser	
Glu Gly Thr Arg 340	-	Ala Gln	Ala Ile 345	Ile As	p Ala	Thr 350	Ala	Gly	
Asn Asp Ala Phe 355	Ser Val	Val Gly 360	Gly Gly	Asp Se	r Ala 365	Ala	Ser	Val	
Arg Val Leu Gly 370	Leu Asn	Glu Asp 375	Gly Phe	Ser Hi 38		Ser	Thr	Gly	
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Ala Ile Leu Ala	Gln 405								
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ataattctag ttag	ctccca a	gttggcat	a ggaggc	Va	g gct l Ala l				115
cac gta ttc gct	cgc gaa	att ctc	gac tcc	Va cgc gg	l Ala l t aac	Glu	Ile acc	Met 5 gtc	
cac gta ttc gct	cgc gaa Arg Glu 10	att ctc Ile Leu gat gac	gac tcc Asp Ser 15	cgc gg Arg Gl	l Ala l t aac y Asn t gtc	Glu cca Pro	acc Thr 20	Met 5 gtc Val gtt	115
cac gta ttc gct His Val Phe Ala gag gca gag gtt Glu Ala Glu Val	cgc gaa Arg Glu 10 ttc ctg Phe Leu	att ctc Ile Leu gat gac Asp Asp	gac tcc Asp Ser 15 ggt tcc Gly Ser 30 cac gag	cgc gg Arg Gl cac gg His Gl	l Ala l t aac y Asn t gtc y Val t gag	Cca Pro gca Ala 35	acc Thr 20 ggt Gly	Met 5 5 gtc Val gtt Val gac	115 163
cac gta ttc gct His Val Phe Ala  gag gca gag gtt Glu Ala Glu Val 25  cca tcc ggc gca Pro Ser Gly Ala	cgc gaa Arg Glu 10  ttc ctg Phe Leu  tcc acc Ser Thr	att ctc Ile Leu  gat gac Asp Asp  ggc gtc Gly Val 45 ggc aag	gac tcc Asp Ser 15 ggt tcc Gly Ser 30 cac gag His Glu ggc gtt	cgc gg Arg Gl cac gg His Gl gct ca Ala Hi	l Ala l t aac y Asn t gtc y Val t gag s Glu 50 g gca s Ala	CCa Pro GCa Ala 35 Ctg Leu	acc Thr 20 ggt Gly cgt Arg	Met 5 gtc Val gtt Val gac Asp	<ul><li>115</li><li>163</li><li>211</li><li>259</li></ul>

_	cgc Arg			-	-	-	_		-		-			_		403
_	tcc Ser	_	_		-		-				-		_	_	_	451
	aag Lys															499
	gga Gly 135			_		-			-		-	_				547
	ggt Gly															595
	gct Ala															643
	gag Glu															691
	acc Thr															739
	cgt Arg 215		-		-			-		-			_	-		787
	acc Thr			-	-		-		-	_	-	_	_			835
	ttc Phe				Ğĺy	Thr		His	Phe	Ğlu	ĞÎy	Gly	Gln	His	Ser	883
	gct Ala															931
	gtc Val															979
	aac Asn 295															1027
	ttc Phe															1075

		gtt aag gtg aac Val Lys Val Asr 335	n Gln Ile Gly	
	e Asp Ala Val	gac atg gct cac Asp Met Ala His 350		
		ggt gag acc gag Gly Glu Thr Glu 365		
		tgt ggc cag ato Cys Gly Gln Ile		
		aag tac aac cag Lys Tyr Asn Glr 400	n Leu Leu Arg	
		gtc tac gca ggt Val Tyr Ala Gly 415	y Arg Ser Ala	
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<211> 425

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<400> 72

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Gly Val Ala Gly Val Pro Ser Gly Ala Ser Thr Gly Val His Glu Ala 35 40 45

His Glu Leu Arg Asp Gly Gly Asp Arg Tyr Leu Gly Lys Gly Val Leu 50 55 60

Lys Ala Val Glu Asn Val Asn Glu Glu Ile Gly Asp Glu Leu Ala Gly 65 70 75 80

Leu Glu Ala Asp Asp Gln Arg Leu Ile Asp Glu Ala Met Ile Lys Leu 85 90 95

Asp Gly Thr Ala Asn Lys Ser Arg Leu Gly Ala Asn Ala Ile Leu Gly 100 105 110

Val Ser Met Ala Val Ala Lys Ala Ala Ala Asp Ser Ala Gly Leu Pro 115 120 125

Leu Phe Arg Tyr Ile Gly Gly Pro Asn Ala His Val Leu Pro Val Pro 130 135 140

Met 1	Met	Asn	Ile	Ile	Thr 150	Gly	Gly	Ala	His	Ala 155	Asp	Ser	Gly	Val	Asp 160
Val	Gln	Glu	Phe	Met 165	Ile	Ala	Pro	Ile	Gly 170	Ala	Glu	Thr	Phe	Ser 175	Glu
Ala	Leu	Arg	Asn 180	Gly	Ala	Glu	Val	Tyr 185	His	Ala	Leu	Lys	Ser 190	Val	Ile
Lys	Glu	Lys 195	Gly	Leu	Ser	Thr	Gly 200	Leu	Gly	Asp	Glu	Gly 205	Gly	Phe	Ala
Pro	Ser 210	Val	Gly	Ser	Thr	Arg 215	Glu	Ala	Leu	Asp	Leu 220	Ile	Val	Glu	Ala
Ile 225	Glu	Lys	Ala	Gly	Phe 230	Thr	Pro	Gly	Lys	Asp 235	Ile	Ala	Leu	Ala	Leu 240
Asp	Val	Ala	Ser	Ser 245	Glu	Phe	Phe	Lys	Asp 250	Gly	Thr	Tyr	His	Phe 255	Glu
Gly	Gly	Gln	His 260	Ser	Ala	Ala	Glu	Met 265	Ala	Asn	Val	Tyr	Ala 270	Glu	Leu
Val .	Asp	Ala 275	Tyr	Pro	Ile	Val	Ser 280	Ile	Glu	Asp	Pro	Leu 285	Gln	Glu	Asp
Asp	Trp 290	Glu	Gly	Tyr	Thr	Asn 295	Leu	Thr	Ala	Thr	Ile 300	Gly	Asp	Lys	Val
Gln 305	Ile	Val	Gly	Asp	Asp 310	Phe	Phe	Val	Thr	Asn 315	Pro	Glu	Arg		Lys 320
Glu	Gly	Ile	Ala	Lys 325	Lys	Ala	Ala	Asn	Ser 330	Ile	Leu	Val	Lys	Val 335	Asn
Gln	Ile	Gly	Thr 340	Leu	Thr	Glu	Thr	Phe 345	Asp	Ala	Val	Asp	Met 350	Ala	His
Arg .	Ala	Gly 355	Tyr	Thr	Ser	Met	Met 360	Ser	His	Arg	Ser	Gly 365	Glu	Thr	Glu
Asp	Thr 370	Thr	Ile	Ala	Asp	Leu 375	Ala	Val	Ala	Leu	Asn 380	Cys	Gly	Gln	Ile
Lys 385	Thr	Gly	Ala	Pro	Ala 390	Arg	Ser	Asp	Arg	Val 395	Ala	Lys	Tyr	Asn	Gln 400
Leu	Leu	Arg	Ile	Glu 405	Gln	Leu	Leu	Gly	Asp 410	Ala	Gly	Val	Tyr	Ala 415	Gly
Arg	Ser	Ala	Phe 420	Pro	Arg	Phe	Gln	Gly 425							

<sup>&</sup>lt;210> 73

<sup>&</sup>lt;211> 1554

<sup>&</sup>lt;212> DNA

<sup>&</sup>lt;213> Corynebacterium glutamicum

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cca gca gat gct gaa ctc gtt cac aag atc atg gac gaa gga ggt cgt

Pro Ala Asp 200	Ala Glu Leu	Val His Lys 205	Ile Met Asp	Glu Glu Gly 210	Arg
cgt gtt cct Arg Val Pro 215	gtg atc gcc Val Ile Ala	aag ctg gaa Lys Leu Glu 220	aag cca gag Lys Pro Glu 225	gct gtc acc Ala Val Thr	tcc 787 Ser
ctc gag cca Leu Glu Pro 230	atc gtg ttg Ile Val Leu 235	gca ttc gac Ala Phe Asp	gcc gtc atg Ala Val Met 240	gtt gct cgt Val Ala Arg	ggt 835 Gly 245
			gag gtt cca Glu Val Pro 255		
cgc gca atc Arg Ala Ile	cag att gcc Gln Ile Ala 265	cgt gag aac Arg Glu Asn 270	gca aag cca Ala Lys Pro	gtt atc gtg Val Ile Val 275	gca 931 Ala
acc cag atg Thr Gln Met 280	ctg gat tcc Leu Asp Ser	atg att gag Met Ile Glu 285	aac tcc cgc Asn Ser Arg	cca acc cgt Pro Thr Arg 290	gcg 979 Ala
gaa gct tct Glu Ala Ser 295	gac gtg gca Asp Val Ala	aac gct gtg Asn Ala Val 300	ctc gat ggc Leu Asp Gly 305	gca gat gct Ala Asp Ala	gtc 1027 Val
atg ctt tct Met Leu Ser 310	ggt gaa act Gly Glu Thr 315	tca gtg ggc Ser Val Gly	aaa gat ccg Lys Asp Pro 320	cac aac gtt His Asn Val	gtg 1075 Val 325
			gct gaa acc Ala Glu Thr 335		Val
			aag cgt ggc Lys Arg Gly		
			aac gct cgt Asn Ala Arg		
ttc acc acc Phe Thr Thr 375	tct ggt gat Ser Gly Asp	acc gca aag Thr Ala Lys 380	cgt gtg gct Arg Val Ala 385	cgt ctg cac Arg Leu His	agc 1267 Ser
_			aat gag gca Asn Glu Ala 400		
			ttc ctg tgt Phe Leu Cys 415		Ser
			gac cgt gct Asp Arg Ala		
			gtt gtt gtt Val Val Val		

440 445 450

cct ggt gtt acc ggt aac acc aac atg att cac gtc cac ctt ctt ggt 1507
Pro Gly Val Thr Gly Asn Thr Asn Met Ile His Val His Leu Leu Gly
455 460 465

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<210> 74

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<212> PRT

<213> Corynebacterium glutamicum

<400> 74

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Val Ala Arg Leu Asn Phe Ser His Gly Asp His Pro Asp His Glu Gln 35 40 45

Asn Tyr Lys Trp Val Arg Glu Ala Ala Glu Lys Thr Gly Arg Ala Val 50 55 60

Gly Ile Leu Ala Asp Leu Gln Gly Pro Lys Ile Arg Leu Gly Arg Phe 65 70 75 80

Thr Asp Gly Ala Thr Val Trp Glu Asn Gly Glu Thr Ile Arg Ile Thr 85 90 .95

Val Asp Asp Val Glu Gly Thr His Asp Arg Val Ser Thr Thr Tyr Lys 100 105 110

Asn Leu Ala Lys Asp Ala Lys Pro Gly Asp Arg Leu Leu Val Asp Asp 115 120 125

Gly Lys Val Gly Leu Val Cys Val Ser Val Glu Gly Asn Asp Val Ile 130 135 140

Cys Glu Val Val Glu Gly Gly Pro Val Ser Asn Asn Lys Gly Val Ser 145 150 155 160

Leu Pro Gly Met Asp Ile Ser Val Pro Ala Leu Ser Glu Lys Asp Ile 165 170 175

Arg Asp Leu Arg Phe Ala Leu Lys Leu Gly Val Asp Phe Ile Ala Leu 180 185 190

Ser Phe Val Arg Ser Pro Ala Asp Ala Glu Leu Val His Lys Ile Met 195 200 205

Asp Glu Glu Gly Arg Arg Val Pro Val Ile Ala Lys Leu Glu Lys Pro 210 215 220

Glu Ala Val Thr Ser Leu Glu Pro Ile Val Leu Ala Phe Asp Ala Val 225 230 235 240

Met Val Ala Arg Gly Asp Leu Gly Val Glu Val Pro Leu Glu Glu Val Pro Leu Val Gln Lys Arg Ala Ile Gln Ile Ala Arg Glu Asn Ala Lys Pro Val Ile Val Ala Thr Gln Met Leu Asp Ser Met Ile Glu Asn Ser Arg Pro Thr Arg Ala Glu Ala Ser Asp Val Ala Asn Ala Val Leu Asp Gly Ala Asp Ala Val Met Leu Ser Gly Glu Thr Ser Val Gly Lys Asp Pro His Asn Val Val Arg Thr Met Ser Arg Ile Val Arg Phe Ala Glu Thr Asp Gly Arg Val Pro Asp Leu Thr His Ile Pro Arg Thr Lys Arg 345 Gly Val Ile Ser Tyr Ser Ala Arg Asp Ile Ala Glu Arg Leu Asn Ala 360 Arg Ala Leu Val Ala Phe Thr Thr Ser Gly Asp Thr Ala Lys Arg Val 375 Ala Arg Leu His Ser His Leu Pro Leu Leu Val Phe Thr Pro Asn Glu 395 390 Ala Val Arg Ser Glu Leu Ala Leu Thr Trp Gly Ala Thr Thr Phe Leu 405 410 Cys Pro Pro Val Ser Asp Thr Asp Asp Met Met Arg Glu Val Asp Arg Ala Leu Leu Ala Met Pro Glu Tyr Asn Lys Gly Asp Met Met Val Val Val Ala Gly Ser Pro Pro Gly Val Thr Gly Asn Thr Asn Met Ile His Val His Leu Leu Gly Asp Asp Thr Arg Ile Ala Lys Leu 470 <210> 75 <211> 1980 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(1957) <223> RXN02675 <400> 75 aagtgtttca ttggaacact tgcgctgcca actttttggt ttacgggcac aatgaaactg 60

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Met Asn Glu Phe Asp cag gac att ctc cag gag atc aag act gaa ctc gac gag tta att cta 163 Gin Asp Ile Leu Gin Glu Ile Lys Thr Glu Leu Asp Glu Leu Ile Leu 15 gaa ctt gat gag gtg aca caa act cac agc gag gcc atc ggg cag gtc 211 Glu Leu Asp Glu Val Thr Gln Thr His Ser Glu Ala Ile Gly Gln Val tcc cca acc cat tac gtt ggt gcc cgc aac ctc atg cat tac gcg cat 259 Ser Pro Thr His Tyr Val Gly Ala Arg Asn Leu Met His Tyr Ala His 307 ctt cgc acc aaa gac ctc cgt ggc ctg cag caa cgc ctc tcc tct gtg Leu Arg Thr Lys Asp Leu Arg Gly Leu Gln Gln Arg Leu Ser Ser Val gga gct acc cgc ttg act acc acc gaa cca gca gtg cag gcc cgc ctc 355 Gly Ala Thr Arg Leu Thr Thr Glu Pro Ala Val Gln Ala Arg Leu ឧ೧ aag gcc gcc cgc aat gtt atc gga gct ttc gca ggt gaa ggc cca ctt 403 Lys Ala Ala Arg Asn Val Ile Gly Ala Phe Ala Gly Glu Gly Pro Leu 95 451 tat cca ccc tca gat gtc gtc gat gcc ttc gaa gat gcc gat gag att Tyr Pro Pro Ser Asp Val Val Asp Ala Phe Glu Asp Ala Asp Glu Ile 105 110 ctc gac gag cac gcc gaa att ctc ctt ggc gaa ccc cta ccg gat act 499 Leu Asp Glu His Ala Glu Ile Leu Leu Gly Glu Pro Leu Pro Asp Thr 125 120 cca tec tgc atc atg gtc acc etg ecc acc gaa gee gee acc gae att 547 Pro Ser Cys Ile Met Val Thr Leu Pro Thr Glu Ala Ala Thr Asp Ile 135 gaa ctt gtc cgt ggc ttc gcc aaa agc ggc atg aat cta gct cgc atc 595 Glu Leu Val Arg Gly Phe Ala Lys Ser Gly Met Asn Leu Ala Arg Ile 160 aac tgt gca cac gac gat gaa acc gtc tgg aag cag atg atc gac aac 643 Asn Cys Ala His Asp Asp Glu Thr Val Trp Lys Gln Met Ile Asp Asn 170 175 qtc cac acc gtt gca gaa gtt ggc cgg gaa atc cgc gtc agc atg 691 Val His Thr Val Ala Glu Glu Val Gly Arg Glu Ile Arg Val Ser Met 190 739 qac etc gec gga eca aaa gta ege ace gge gaa ate gee eca gge gea Asp Leu Ala Gly Pro Lys Val Arg Thr Gly Glu Ile Ala Pro Gly Ala 205 787 qaa qta qqt cqc qca cqa qta acc cgc gac gaa acc gga aaa gta ctg Glu Val Gly Arg Ala Arg Val Thr Arg Asp Glu Thr Gly Lys Val Leu 835 acg ccc gca aaa ctg tgg atc acc gcc cac ggc tcc gaa cca gtc cca Thr Pro Ala Lys Leu Trp Ile Thr Ala His Gly Ser Glu Pro Val Pro

230	235	240	245
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cca gaa tgg ttc gac Pro Glu Trp Phe Asp 265	aaa cta gaa atc gg Lys Leu Glu Ile Gl 270	gc agc gtc atc aac gtc y Ser Val Ile Asn Val 275	cca 931 Pro
		ce gtg acc agg gtt ttt or Val Thr Arg Val Phe 290	
		aa gcc tac atc tcc aac vs Ala Tyr Ile Ser Asn 305	
		cc cgg gtc tac ggc atc er Arg Val Tyr Gly Ile 320	
	Ile Asn Leu Lys Va	cc ggc gac cgc ctc atc al Gly Asp Arg Leu Ile 35 340	
		ce etc gga tec gge ege er Leu Gly Ser Gly Arg 355	
		ca gtc gat gca att aaa La Val Asp Ala Ile Lys 370	
		ce ate gee gea gte tge La Ile Ala Ala Val Cys 385	
		nc gac gta gaa ttg gaa sn Asp Val Glu Leu Glu 400	
	Gln Gly Val Asn Le	g gcc gca tac aag gga eu Ala Ala Tyr Lys Gly .5 420	
aac ctc cca gac tcc Asn Leu Pro Asp Ser 425	gaa ctt cca ctc cc Glu Leu Pro Leu Pr 430	ca agc ctc act gaa gaa co Ser Leu Thr Glu Glu 435	gac 1411 Asp
		ac gcc gac atc gca gcc yr Ala Asp Ile Ala Ala 450	
		aa tac ctc ctc caa gca .u Tyr Leu Leu Gln Ala 465	
		aa cgc ctt ggc ctc gtc .u Arg Leu Gly Leu Val 480	

Lys Ile Glu	acc atc Thr Ile 490											1603
acc ggc atg Thr Gly Met												1651
ctc gcc gtc Leu Ala Val 520												1699
atc atg gcc Ile Met Ala 535			Āla									1747
caa gtc ctg Gln Val Leu 550												1795
atc acc gac Ile Thr Asp												1843
aag gga cca Lys Gly Pro												1891
cgc aaa ctt Arg Lys Leu												1939
600			003					010				
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Gly	Glu	Gly	Pro 100	Leu	Tyr	Pro	Pro	Ser 105	Asp	Val	Val	Asp	Ala 110	Phe	Glu
Asp	Ala	Asp 115	Glu	Ile	Leu	Asp	Glu 120	His	Ala	Glu	Ile	Leu 125	Leu	Gly	Glu
Pro	Leu 130	Pro	Asp	Thr	Pro	Ser 135	Cys	Ile	Met	Val	Thr 140	Leu	Pro	Thr	Glu
Ala 145	Ala	Thr	Asp	Ile	Glu 150	Leu	Val	Arg	Gly	Phe 155	Ala	Lys	Ser	Gly	Met 160
Asn	Leu	Ala	Arg	Ile 165	Asn	Cys	Ala	His	Asp 170	Asp	Glu	Thr	Val	Trp 175	Lys
Gln	Met	Ile	Asp 180	Asn	Val	His	Thr	Val 185	Ala	Glu	Glu	Val	Gly 190	Arg	Glu
Ile	Arg	Val 195	Ser	Met	Asp	Leu	Ala 200	Gly	Pro	Lys	Val	Arg 205	Thr	Gly	Glu
Ile	Ala 210	Pro	Gly	Ala	Glu	Val 215	Gly	Arg	Ala	Arg	Val 220	Thr	Arg	Asp	Glu
Thr 225	Gly	Lys	Val	Leu	Thr 230	Pro	Ala	Lys	Leu	Trp 235	Ile	Thr	Ala	His	Gly 240
Ser	Glu	Pro	Val	Pro 245	Ala	Pro	Glu	Ser	Leu 250	Pro	Gly	Arg	Pro	Ala 255	Leu
Pro	Ile	Glu	Val 260	Thr	Pro	Glu	Trp	Phe 265	Asp	Lys	Leu	Glu	Ile 270	Gly	Ser
Val	Ile	Asn 275	Val	Pro	Asp	Thr	Arg 280	Gly	Ser	Arg	Arg	Ala 285	Phe	Thr	Val
Thr	Arg 290	Val	Phe	Asp	Gly	Ala 295	Val	Leu	Ala	Glu	Gly 300	Pro	Gln	Lys	Ala
Tyr 305	Ile	Ser	Asn	Gly	Thr 310	Leu	Leu	Glu	His	Asn 315	Tyr	Asp	Arg	Ser	Arg 320
Val	Tyr	Gly	Ile	Pro 325	Ala	Val	Val	Gln	Arg 330	Ile	Asn	Leu	Lys	Val 335	Gly
Asp	Arg	Leu	Ile 340	Leu	Thr	Asp	Glu	Glu 345	Leu	Thr	Tyr	Asp	Pro 350	Ser	Leu
Gly	Ser	Gly 355	Arg	Thr	Pro	Arg	Ile 360	Ser	Cys	Thr	Leu	Pro 365	Gln	Ala	Val
Asp	Ala 370	Ile	Lys	Val	Gly	His 375	Arg	Val	Leu	Phe	Asp 380	Asp	Gly	Ala	Ile
Ala 385	Ala	Val	Cys	Ile	Asp 390	Lys	Thr	Ser	Thr	Ala 395	Asp	Gly	His	Asn	Asp 400
Val	Glu	Leu	Glu	Val 405	Thr	His	Ala	Arg	Pro 410	Gln	Gly	Val	Asn	Leu 415	Ala

Ala Tyr Lys Gly Ile Asn Leu Pro Asp Ser Glu Leu Pro Leu Pro Ser 425 Leu Thr Glu Glu Asp Leu Gln His Leu Arg Phe Val Val Lys Tyr Ala Asp Ile Ala Ala Ile Ser Phe Ile Arg Asn Val Ala Asp Val Glu Tyr Leu Leu Gln Ala Leu Ala Asp Ile Gly Asp Pro Val Ala Val Glu Arg 475 Leu Gly Leu Val Leu Lys Ile Glu Thr Ile Pro Gly Tyr Glu Gly Leu Ala Gln Ile Leu Leu Thr Gly Met Arg His Glu Asn Phe Gly Ile Met Ile Ala Arg Gly Asp Leu Ala Val Glu Leu Gly Phe Asp Arg Met Ala Glu Val Pro Gln Leu Ile Met Ala Leu Ala Glu Ala Ala His Val Pro 535 Thr Ile Leu Ala Thr Gln Val Leu Glu Asn Met Ala Lys Asn Gly Leu 550 555 Pro Ser Arg Ala Glu Ile Thr Asp Ala Ala Met Ala Leu Arg Ala Glu 570 Cys Val Met Leu Asn Lys Gly Pro His Ile Asn Asp Ala Ile Lys Val Leu Thr Glu Met Ser Arg Lys Leu Gly Ala Ser Gln Arg Lys Ser Arg 595 Leu Leu Leu Arg Lys Val Lys Ser Trp Glu Glu 610 <210> 77 <211> 386 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (1)..(363) <223> FRXA02675 <400> 77 atc ctc atg acc ggc atg cgc cac gaa aac ttc ggc atc atg atc gcc 48 Ile Leu Met Thr Gly Met Arg His Glu Asn Phe Gly Ile Met Ile Ala cgc gga gac etc gec gtc gaa etc ggc ttc gac egc atg gea gaa gtc 96 Arg Gly Asp Leu Ala Val Glu Leu Gly Phe Asp Arg Met Ala Glu Val ccc caa ctg atc atg gcc ctt gca gaa gcc gcc cac gtc cca acc atc Pro Gln Leu Ile Met Ala Leu Ala Glu Ala Ala His Val Pro Thr Ile

40 35 45 ttg gcc acc caa gtc ctg gaa aac atg gcc aaa aac gga ctc cca tct 192 Leu Ala Thr Gln Val Leu Glu Asn Met Ala Lys Asn Gly Leu Pro Ser cgc gca gaa atc acc gac gca gca atg gca ctt cgc gct gaa tgc gtc 240 Arg Ala Glu Ile Thr Asp Ala Ala Met Ala Leu Arg Ala Glu Cys Val atg ctg aac aag gga cca cac atc aac gac gcc atc aag gtc ctc acc Met Leu Asn Lys Gly Pro His Ile Asn Asp Ala Ile Lys Val Leu Thr 85 gaa atg agc cgc aaa ctt ggt gca tcc caa cga aag agt agg ctg ctg 336 Glu Met Ser Arg Lys Leu Gly Ala Ser Gln Arg Lys Ser Arg Leu Leu 110 ctg cgc aag gtg aag agc tgg gaa gag taactcacaa aggcgattgg 383 Leu Arg Lys Val Lys Ser Trp Glu Glu 386 cqt <210> 78 <211> 121 <212> PRT <213> Corynebacterium glutamicum <400> 78 Ile Leu Met Thr Gly Met Arg His Glu Asn Phe Gly Ile Met Ile Ala Arg Gly Asp Leu Ala Val Glu Leu Gly Phe Asp Arg Met Ala Glu Val Pro Gln Leu Ile Met Ala Leu Ala Glu Ala Ala His Val Pro Thr Ile Leu Ala Thr Gln Val Leu Glu Asn Met Ala Lys Asn Gly Leu Pro Ser Arg Ala Glu Ile Thr Asp Ala Ala Met Ala Leu Arg Ala Glu Cys Val Met Leu Asn Lys Gly Pro His Ile Asn Asp Ala Ile Lys Val Leu Thr Glu Met Ser Arg Lys Leu Gly Ala Ser Gln Arg Lys Ser Arg Leu Leu 100 Leu Arg Lys Val Lys Ser Trp Glu Glu 115 <210> 79 <211> 1522

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acg ccc gca aaa Thr Pro Ala Lys 230	ctg tgg atc Leu Trp Ile 235	acc gcc cac Thr Ala His	ggc tcc gaa Gly Ser Glu 240	cca gtc Pro Val	cca 835 Pro 245
gcc ccc gaa agc Ala Pro Glu Ser					
cca gaa tgg ttc Pro Glu Trp Phe 265	Asp Lys Leu	•		-	
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ggc gcg gtc ctc Gly Ala Val Leu 295		Pro Gln Lys			
acc ctc ctg gaa Thr Leu Leu Glu 310		-			
gcc gta gtt cag Ala Val Val Gln					
acc gac gaa gaa Thr Asp Glu Glu 345		-			
cca cgc atc agc Pro Arg Ile Ser 360					
ggg cac cgc gtg Gly His Arg Val 375	_		- :	-	
gac aag acc tcc Asp Lys Thr Ser 390					_
acc cac gcc cgc Thr His Ala Arg					
aac ctc cca gac Asn Leu Pro Asp 425	-		-		-
ctc caa cac ctg Leu Gln His Leu					

440 445 450

tcc ttc atc cga aac gtc gcc gac gtg gaa tac ctc ctc caa gca ctc 1507 Ser Phe Ile Arg Asn Val Ala Asp Val Glu Tyr Leu Leu Gln Ala Leu

gcc gac atc gga gat 1522 Ala Asp Ile Gly Asp

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<213> Corynebacterium glutamicum

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Ala Ile Gly Gln Val Ser Pro Thr His Tyr Val Gly Ala Arg Asn Leu
35 45

Met His Tyr Ala His Leu Arg Thr Lys Asp Leu Arg Gly Leu Gln Gln 50 60

Arg Leu Ser Ser Val Gly Ala Thr Arg Leu Thr Thr Glu Pro Ala 65 70 75 80

Val Gln Ala Arg Leu Lys Ala Ala Arg Asn Val Ile Gly Ala Phe Ala 85 90 95

Gly Glu Gly Pro Leu Tyr Pro Pro Ser Asp Val Val Asp Ala Phe Glu 100 105 110

Asp Ala Asp Glu Ile Leu Asp Glu His Ala Glu Ile Leu Leu Gly Glu 115 120 125

Pro Leu Pro Asp Thr Pro Ser Cys Ile Met Val Thr Leu Pro Thr Glu 130 135 140

Ala Ala Thr Asp Ile Glu Leu Val Arg Gly Phe Ala Lys Ser Gly Met 145 150 155 160

Asn Leu Ala Arg Ile Asn Cys Ala His Asp Asp Glu Thr Val Trp Lys 165 170 175

Gln Met Ile Asp Asn Val His Thr Val Ala Glu Glu Val Gly Arg Glu 180 185 190

Ile Arg Val Ser Met Asp Leu Ala Gly Pro Lys Val Arg Thr Gly Glu 195 200 205

Ile Ala Pro Gly Ala Glu Val Gly Arg Ala Arg Val Thr Arg Asp Glu 210 215 220

Thr Gly Lys Val Leu Thr Pro Ala Lys Leu Trp Ile Thr Ala His Gly 225 230 235 240

Ser Glu Pro Val Pro Ala Pro Glu Ser Leu Pro Gly Arg Pro Ala Leu Pro Ile Glu Val Thr Pro Glu Trp Phe Asp Lys Leu Glu Ile Gly Ser Val Ile Asn Val Pro Asp Thr Arg Gly Ser Arg Arg Ala Phe Thr Val 280 Thr Arg Val Phe Asp Gly Ala Val Leu Ala Glu Gly Pro Gln Lys Ala Tyr Ile Ser Asn Gly Thr Leu Leu Glu His Asn Tyr Asp Arg Ser Arg 310 315 Val Tyr Gly Ile Pro Ala Val Val Gln Arg Ile Asn Leu Lys Val Gly Asp Arg Leu Ile Leu Thr Asp Glu Glu Leu Thr Tyr Asp Pro Ser Leu Gly Ser Gly Arg Thr Pro Arg Ile Ser Cys Thr Leu Pro Gln Ala Val 360 Asp Ala Ile Lys Val Gly His Arg Val Leu Phe Asp Asp Gly Ala Ile 375 380 Ala Ala Val Cys Ile Asp Lys Thr Ser Thr Ala Asp Gly His Asn Asp 390 Val Glu Leu Glu Val Thr His Ala Arg Pro Gln Gly Val Asn Leu Ala Ala Tyr Lys Gly Ile Asn Leu Pro Asp Ser Glu Leu Pro Leu Pro Ser 425 Leu Thr Glu Glu Asp Leu Gln His Leu Arg Phe Val Val Lys Tyr Ala Asp Ile Ala Ala Ile Ser Phe Ile Arg Asn Val Ala Asp Val Glu Tyr Leu Leu Gln Ala Leu Ala Asp Ile Gly Asp 470 <210> 81 <211> 2022 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(1999) <223> RXA00682 <400> 81 ataggcacct tcgatttcag ctcaatcacc gtcgcaatga ccggcacgaa gtaaaaccac 60

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Met Ala Asn Lys Ser ttc ccc aag ccc tcc gat ctt cca gtg ccc aag ggc gct gaa ggt tgg 163 Phe Pro Lys Pro Ser Asp Leu Pro Val Pro Lys Gly Ala Glu Gly Trp qaa qat ctq tac ccq tac tac ctc gtt ttc caa gac aag ctc atg gat 211 Glu Asp Leu Tyr Pro Tyr Tyr Leu Val Phe Gln Asp Lys Leu Met Asp caa gag aat gag aaa ttc tgg ttc tgc gat tca cag cac tgg cca act 259 Gln Glu Asn Glu Lys Phe Trp Phe Cys Asp Ser Gln His Trp Pro Thr gtg ttc aag cct ttt gaa act atc ggt ggt gaa ttc gct gta aag tgc 307 Val Phe Lys Pro Phe Glu Thr Ile Gly Gly Glu Phe Ala Val Lys Cys ctc ggc caa tac aac gct cgg cat ttg atg atc ccg aat gcc aat ggc 355 Leu Gly Gln Tyr Asn Ala Arg His Leu Met Ile Pro Asn Ala Asn Gly atc gag ttc cgc gtg cat ctg gga tac ctc tat atg tcc cct att cca 403 Ile Glu Phe Arg Val His Leu Gly Tyr Leu Tyr Met Ser Pro Ile Pro 95 gtg cct gaa gat cag att gcg gaa cgc gtc ccc atg ttc cag gaa cgc 451 Val Pro Glu Asp Gln Ile Ala Glu Arg Val Pro Met Phe Gln Glu Arg 105 110 atc acg cac tac ttc caa aac tgg gag cca atg ctg gca aat tgg aag 499 Ile Thr His Tyr Phe Gln Asn Trp Glu Pro Met Leu Ala Asn Trp Lys 120 125 gaq cqa gta tta gga acc atc aat gag ctg gaa tct cta gaa ttc aag 547 Glu Arg Val Leu Gly Thr Ile Asn Glu Leu Glu Ser Leu Glu Phe Lys cca ctg cct gac tac gtg cct atc gat gat att gtc tcc gga aaa gcc 595 Pro Leu Pro Asp Tyr Val Pro Ile Asp Asp Ile Val Ser Gly Lys Ala aaa qac ggc acc gaa gta ctc atg gaa aac ttc gat cgg ctc att cag 643 Lys Asp Gly Thr Glu Val Leu Met Glu Asn Phe Asp Arg Leu Ile Gln 170 175 ctc gcc tac caa aac tgg caa tac cac ttt gag ttc ctc aac ttg ggt 691 Leu Ala Tyr Gln Asn Trp Gln Tyr His Phe Glu Phe Leu Asn Leu Gly 190 tac atc gct tac cta gat ttc ttc aat ttc tgc aag gaa gtc ttc cca 739 Tyr Ile Ala Tyr Leu Asp Phe Phe Asn Phe Cys Lys Glu Val Phe Pro 205 787 gat atc cct gat caa tca att tcg atg atg gtt cag ggc gtg gat atg Asp Ile Pro Asp Gln Ser Ile Ser Met Met Val Gln Gly Val Asp Met 220 835 gag ctg ttc cgc ccc gat gat gaa cta aag att ctg gca cag cta gcg Glu Leu Phe Arg Pro Asp Glu Leu Lys Ile Leu Ala Gln Leu Ala

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		g ttc aac ttc acc gtc p Phe Asn Phe Thr Val 290	
		g atc gag cac ctg gaa p Ile Glu His Leu Glu 305	
		c cgc cta gat gaa ggc g Arg Leu Asp Glu Gly 320	
		c gca gaa aag gaa cgc e Ala Glu Lys Glu Arg 5 340	
		a gaa caa ctc gcg cag y Glu Gln Leu Ala Gln 355	
		a tac ccc tat gtg gaa a Tyr Pro Tyr Val Glu 370	
		g tca gta ttt tgg cgc et Ser Val Phe Trp Arg 385	
		te tac ggt ttc tgg gag y Tyr Gly Phe Trp Glu 400	
		t gaa gtc cgc gat gtc r Glu Val Arg Asp Val 5 420	
		c gca ccc ggt ggt cca y Ala Pro Gly Gly Pro 435	
	, , , , ,	g cga aga aaa gca att u Arg Arg Lys Ala Ile 450	-
		a gct ctt aac act cct o Ala Leu Asn Thr Pro 465	
	_	g ctc tgg gga atc acc t Leu Trp Gly Ile Thr 480	

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acc ctt aaa Thr Leu Lys	ggc atg Gly Met 505	gct (	gca Ala	tcc Ser	cct Pro 510	ggt Gly	gtg Val	gtg Val	gaa Glu	ggc Gly 515	tac Tyr	gct Ala	1651
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atc ctc gtt Ile Leu Val 535	gcc cct Ala Pro	Val :	aca Thr 540	gca Ala	cct Pro	tct Ser	tgg Trp	ggc Gly 545	cca Pro	atc Ile	ttt Phe	ggc Gly	1747
aaa atc aag Lys Ile Lys 550	gca aca Ala Thr	gtc a Val 5 555	act Thr	gat Asp	att Ile	ggt Gly	ggc Gly 560	atg Met	atg Met	agc Ser	cat His	gct Ala 565	1795
gcg atc gtg Ala Ile Val	tgc cgc Cys Arg 570	gaa d Glu d	tac Tyr	ggc Gly	ttg Leu	ccg Pro 575	gct Ala	gtt Val	act Thr	gga Gly	act Thr 580	ggc Gly	1843
gct gca tcc Ala Ala Ser	acc acc Thr Thr 585	atc : Ile :	aaa Lys	acc Thr	ggc Gly 590	gat Asp	tac Tyr	ctc Leu	aag Lys	gtc Val 595	gat Asp	gga Gly	1891
acc aag ggc Thr Lys Gly 600	aag gtt Lys Val	gtc ( Val	Ile	gtt Val 605	gat Asp	cca Pro	gat Asp	gcg Ala	cca Pro 610	cgc Arg	atc .Ile	gaa Glu	1939
gga ccc ggc Gly Pro Gly 615	gcg cac Ala His	Ser .	cat His 620	gcg Ala	cac His	tca Ser	gta Val	gca Ala 625	gca Ala	cat His	Gly	gtg Val	1987
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	20 Met Asp	Gln	Glu	40	Glu				45	Cys			

65					70					75					80
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Met	Ser	Pro	Ile 100	Pro	Val	Pro	Glu	Asp 105	Gln	Ile	Ala	Glu	Arg 110	Val	Pro
Met	Phe	Gln 115	Glu	Arg	Ile	Thr	His 120	Tyr	Phe	Gln	Asn	Trp 125	Glu	Pro	Met
Leu	Ala 130	Asn	Trp	Lys	Glu	Arg 135	Val	Leu	Gly	Thr	Ile 140	Asn	Glu	Leu	Glu
Ser 145	Leu	Glu	Phe	Lys	Pro 150	Leu	Pro	Asp	Tyr	Val 155	Pro	Ile	Asp	Asp	Ile 160
Val	Ser	Gly	Lys	Ala 165	Lys	Asp	Gly	Thr	Glu 170	Val	Leu	Met	Glu	Asn 175	Phe
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Phe	Leu	Asn 195	Leu	Gly	Tyr	Ile	Ala 200	Tyr	Leu	Asp	Phe	Phe 205	Asn	Phe	Cys
Lys	Glu 210	Val	Phe	Pro	Asp	Ile 215	Pro	Asp	Gln	Ser	Ile 220	Ser	Met	Met	Val
Gln 225	Gly	Val	Asp	Met	Glu 230	Leu	Phe	Arg	Pro	Asp 235	Asp	Glu	Leu	Lys	11e 240
Leu	Ala	Gln	Leu	Ala 245	Val	Asp	Leu	Gly	Leu 250	Gln	Thr	His	Phe	Ala 255	Asn
Pro	Asp	Asp	Pro 260	Gln	Ala	Thr	Leu	Ala 265	Ala	Ile	Ala	Lys	Ala 270	Glu	Gly
Gly	Ala	Thr 275	Trp	Ile	Ala	Arg	Trp 280	Glu	Glu	Ala	Gln	Asp 285	Pro	Trp	Phe
Asn	Phe 290	Thr	Val	Gly	Asn	Gly 295	Phe	Tyr	Gly	His	Asp 300	Lys	Tyr	Trp	Ile
Glu 305	His	Leu	Glu	Leu	Pro 310	Leu	Gly	Tyr	Ile	Ala 315	Asp	Tyr	Ile	Arg	Arg 320
Leu	Asp	Glu	Gly	Gln 325	Thr	Ile	Ser	Arg	Pro 330	Lys	Asp	Glu	Leu	Ile 335	Ala
Glu	Lys	Glu	Arg 340	Val	Val	Glu	Glu	Tyr 345	Arg	Asp	Leu	Leu	Asp 350	Gly	Glu
Gln	Leu	Ala 355	Gln	Phe	Asp	Ala	Lys 360	Cys	Gly	Leu	Ala	Ala 365	Thr	Ala	Tyr
Pro	Tyr 370	Val	Glu	Asn	His	Asn 375	Phe	Tyr	Ile	Glu	His 380	Trp	Thr	Met	Ser
Val 385	Phe	Trp	Arg	Lys	Val 390	Arg	Glu	Leu	Ser	Arg 395	Thr	Leu	Gln	Gly	Tyr 400

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gaa Glu	gta Val	ctc Leu	ggt Gly 25	ggc Gly	aag Lys	ggc Gly	gct Ala	tca Ser 30	cta Leu	gtc Val	acc Thr	atg Met	aca Thr 35	gat Asp	gct Ala	211
gga Gly																259
gaa Glu																307
aac Asn 70																355
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ttc Phe																451
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ttg Leu 150																595
						tac Tyr										643
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gca Ala																739
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gaa Glu	gtg Val	aca Thr	cca Pro	gac Asp	aat Asn	atc Ile	ttg Leu	ctg Leu	gac Asp	aag Lys	atc Ile	acg Thr	ctg Leu	cag Gln	gtt Val	835

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		eg ctc gct gtg gca caa et Leu Ala Val Ala Glr 290	
		gc cca caa gat atc gaa ys Pro Gln Asp Ile Glu 305	
		aa aac ctt ctg tta ttg lu Asn Leu Leu Leu Leu 320	
	His Ser Asn Gl	gt gtg aag aag gaa acc ly Val Lys Lys Glu Thi 335	
		cc ttc gat ttc agc tca nr Phe Asp Phe Ser Sen 50 355	: Ile Thr
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Glu Arg Cys Gly Gly Asp Val Pro Val Ala Val Arg Ser Ser Ala Thr Ala Glu Asp Leu Pro Asp Ala Ser Phe Ala Gly Gln Gln Asp Thr Tyr Leu Trp Gln Val Gly Leu Ser Ala Val Thr Glu His Ile Arg Lys Cys Trp Ala Ser Leu Phe Thr Ser Arg Ala Ile Ile Tyr Arg Leu Lys Asn Asn Ile Pro Asn Glu Gly Leu Ser Met Ala Val Val Gln Lys Met Val Asn Ser Arg Val Ala Gly Val Ala Ile Thr Met Asn Pro Ser Asn 200 Gly Asp Arg Ser Lys Ile Thr Ile Asp Ser Ser Trp Gly Val Gly Glu 215 220 Met Val Val Ser Gly Glu Val Thr Pro Asp Asn Ile Leu Leu Asp Lys 230 235 Ile Thr Leu Gln Val Val Ser Glu His Ile Gly Ser Lys His Ala Glu 250 245 Leu Ile Pro Asp Ala Thr Ser Gly Ser Leu Val Glu Lys Pro Val Asp Glu Glu Arg Ala Asn Arg Arg Ser Leu Thr Asp Glu Glu Met Leu Ala Val Ala Gln Met Ala Lys Arg Ala Glu Lys His Tyr Lys Cys Pro Gln Asp Ile Glu Trp Ala Leu Asp Ala Asp Leu Pro Asp Gly Glu Asn Leu Leu Leu Gln Ser Arg Pro Glu Thr Ile His Ser Asn Gly Val Lys 330 Lys Glu Thr Pro Thr Pro Gln Ala Ala Lys Thr Ile Gly Thr Phe Asp Phe Ser Ser Ile Thr Val Ala Met Thr Gly Thr Lys 360 <210> 85

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agc aag gcg Ser Lys Ala 550	-	_		_									1795
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Asp	Ile	Ala	Lys	Glu 165	Asp	Ala	Gly	Asp	Gly 170	Thr	Tyr	Ser	Asn	Ser 175	Thr
Ile	Ser	Ser	Gly 180	Thr	Pro	Val	Val	Phe 185	Pro	Asp	Pro	Thr	Glu 190	Ala	Ala
Ala	Leu	Val 195	Glu	Ala	Ile	Asn	Asn 200	Ala	Lys	Ser	Val	Thr 205	Leu	Phe	Cys
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Gly	Ala	Cys	Val 260	Asp	Ala	Ser	Asn	Glu 265	Ala	Asp	Leu	Leu	Ile 270	Leu	Leu
Gly	Thr	Asp 275	Phe	Pro	Tyr	Ser	Asp 280	Phe	Leu	Pro	Lys	Asp 285	Asn	Val	Ala
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Ala	His	Glu	Arg 340	Lys	Leu	Ser	Ser	Val 345	Val	Glu	Thr	Tyr	Thr 350	His	Asn
Val	Glu	Lys 355	His	Val	Pro	Ile	His 360	Pro	Glu	Tyr	Val	Ala 365	Ser	Ile	Leu
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Cys 385	Asn	Val	Trp	His	Ala 390	Arg	Tyr	Ile	Glu	Asn 395	Pro	Glu	Gly	Thr	Arg 400
Asp	Phe	Val	Gly	Ser 405	Phe	Arg	His	Gly	Thr 410	Met	Ala	Asn	Ala	Leu 415	Pro
His	Ala	Ile	Gly 420	Ala	Gln	Ser	Val	Asp 425	Arg	Asn	Arg	Gln	Val 430	Ile	Ala
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354

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att go Ile G															450
gaa te Glu C	gc t ys S	ct gg er Gl 12	y Tyr	tgc Cys	gag Glu	atg Met	gtg Val 130	aat Asn	ggt Gly	ggt Gly	gag Glu	cag Gln 135	ggt Gly	gaa Glu	498
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Val A	_	1 - 17-		Gl n	60~	n									
		35	l Arg	GIII	Ser	40	Ile	Glu	Trp	Val	His 45	Val	Arg	Asn	
Glu G		35				40					45				
	lu A 50	35 la Al	a Ala	Phe	Ala 55	40 Ala	Gly	Ala	Glu	Ser 60	45 Leu	Ile	Thr	Gly	
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Glu L 65 Ile G Ile A Glu T	eu A ln G la S hr H	35 la Al la Va ly Le er Hi 10 is Pr 15	A Ala Cys Tyr 85 S Ile	Phe Ala 70 Asp Pro	Ala 55 Ala Ser Ser	40 Ala Ser His Ala Phe 120	Gly Cys Arg Gln 105 Lys	Ala Gly Asn 90 Ile Glu	Glu Pro 75 Gly Gly Cys	Ser 60 Gly Ala Ser	45 Leu Asn Lys Thr Gly 125	Thr Val Phe 110	Thr His Leu 95 Phe Cys	Gly Leu 80 Ala Gln	

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PCT/IB00/00943 WO 01/00844

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120

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	cgt Arg 55															307
	tac Tyr															355
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	cgc Arg															451
	ggt Gly															499
	gga Gly 135															547
	cgc Arg															595
	gat Asp															643
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	ggt Gly 215															787
	gct Ala															835
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954

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Ser Thr Asn Glu Gly Val Glu Thr Tyr Asp Pro Ser Phe Ala Tyr Glu 35 40 45

Ile Ala His Leu Val His Arg Gly Ile Asp Arg Met Tyr Gly Pro Gly 50 55 60

Lys Gly Glu Asp Val Ile Tyr Tyr Ile Thr Ile Tyr Asn Glu Pro Thr 65 70 75 80

Pro Gln Pro Ala Glu Pro Glu Gly Leu Asp Val Glu Gly Leu His Lys 85 90 95

Gly Ile Tyr Leu Tyr Ser Arg Gly Glu Gly Thr Gly His Glu Ala Asn  $100 \hspace{1cm} 105 \hspace{1cm} 110$ 

Ile Leu Ala Ser Gly Val Gly Met Gln Trp Ala Leu Lys Ala Ala Ser 115 120 125

Ile Leu Glu Ala Asp Tyr Gly Val Arg Ala Asn Ile Tyr Ser Ala Thr 130 135 140

Ser Trp Val Asn Leu Ala Arg Asp Gly Ala Ala Arg Asn Lys Ala Gln 145 150 155 160

Leu Arg Asn Pro Gly Ala Asp Ala Gly Glu Ala Phe Val Thr Thr Gln
165 170 175

Leu Lys Gln Thr Ser Gly Pro Tyr Val Ala Val Ser Asp Phe Ser Thr 180 185 190

Asp Leu Pro Asn Gln Ile Arg Glu Trp Val Pro Gly Asp Tyr Thr Val 195 200 205

Leu Gly Ala Asp Gly Phe Gly Phe Ser Asp Thr Arg Pro Ala Ala Arg 210 215 220

Arg Phe Phe Asn Ile Asp Ala Glu Ser Ile Val Val Ala Val Leu Asn 225 230 235 240

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Trp Val Pro Gly Asp Tyr Thr Val Leu Gly Ala Asp Gly Phe Gly Phe
tet gat acc ege eca get get egt ege tte tte aac ate gae get gag
Ser Asp Thr Arg Pro Ala Ala Arg Arg Phe Phe Asn Ile Asp Ala Glu
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tcc att gtt gtt gca gtg ctg aac tcc ctg gca cgc gaa ggc aag atc
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     50
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Asp Val Ser Val Ala Ala Gln Ala Ala Glu Lys Phe Lys Leu Asp Asp
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65
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Ser Asp Thr Arg Pro Ala Ala Arg Arg Phe Phe Asn Ile Asp Ala Glu
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Asp Val Ser Val Ala Ala Gln Ala Ala Glu Lys Phe Lys Leu Asp Asp
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					gtg Val											691
					cca Pro											739
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_	-				gtt Val 235	-	-		_			_	-	-	_	835
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Ile Ala His Leu Val His Arg Gly Ile Asp Arg Met Tyr Gly Pro Gly 50 55 60

Lys Gly Glu Asp Val Ile Tyr Tyr Ile Thr Ile Tyr Asn Glu Pro Thr 65 70 75 80

Pro Gln Pro Ala Glu Pro Glu Gly Leu Asp Val Glu Gly Leu His Lys
85 90 95

Gly Ile Tyr Leu Tyr Ser Arg Gly Glu Gly Thr Gly His Glu Ala Asn 100 105 110

Ile Leu Ala Ser Gly Val Gly Met Gln Trp Ala Leu Lys Ala Ala Ser 115 120 125

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Leu Lys Gln Thr Ser Gly Pro Tyr Val Ala Val 180 185	Ser Asp Phe Ser Thr 190												
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													atg Met			403
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Pro	Glu	Glu 35	Thr	Asn	Glu	Trp	Met 40	Asp	Ser	Leu	Asp	Gly 45	Leu	Leu	Gln	
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Lys Leu Gly Gly Lys Pro Ser Asp Asp Ser Asn Phe Ala Met Ile Arg
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Glu Trp Met Asp Ser Leu Asp Gly Leu Leu Gln Glu Ser Ser Pro Glu
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Arg Val Ser Leu Pro Pro Met Thr Ser Thr Asp Tyr Val Asn Thr Ile
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                                          80
                                                                   403
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Pro Thr Ser Met Glu Pro Glu Phe Pro Gly Asp Glu Glu Met Glu Lys
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Arg Tyr Arg Arg Trp Ile Arg Trp Asn Ala Ala Ile Met Val His Arg
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get cag ega eca gge ate gge gte gge gga cae att tee aet tae gea
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145

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					gtt Val											432
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-		_	-	-	atc Ile		_			_				-		624
					gcc Ala											672
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					aac Asn											768
_	~	_	_	-	ttc Phe	-	-	_	-						_	816
					gat Asp											864
					atc Ile											912
					gag Glu 310											960
		-	-	-	ctt Leu	-		-	_	_						1008
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Asn Thr Lys Ile Ile Gln Glu Leu Glu Ser Phe Phe Arg Gly Ala Gly
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Met	Thr	Asp 195	Glu	Glu	Ile	Trp	Lys 200	Leu	Pro	Arg	Gly	Gly 205	His	Asp	Tyr
Arg	Lys 210	Val	Tyr	Ala	Ala	Tyr 215	Lys	Arg	Ala	Leu	Glu 220	Thr	Lys	Asp	Arg
Pro 225	Thr	Val	Ile	Leu	Ala 230	His	Thr	Ile	Lys	Gly 235	Tyr	Gly	Leu	Gly	His 240
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Glu	Gln	Leu 275	Glu	Lys	Asp	Pro	Tyr 280	Leu	Pro	Pro	Tyr	Tyr 285	His	Pro	Gly
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Pro	Pro	Leu	Asp	Lys 325	Leu	Arg	Ser	Val	Arg 330	Lys	Gly	Ser	Gly	Lys 335	Gln
Gln	Ile	Ala	Thr 340	Thr	Met	Ala	Thr	Val 345	Arg	Thr	Phe	Lys	Glu 350	Leu	Met
Arg	Asp	Lys 355	Gly	Leu	Ala	Asp	Arg 360	Leu	Val	Pro	Ile	Ile 365	Pro	Asp	Glu
Ala	Arg 370	Thr	Phe	Gly	Leu	Asp 375	Ser	Trp	Phe	Pro	Thr 380	Leu	Lys	Ile	Tyr
Asn 385	Pro	His	Gly	Gln	Asn 390	Tyr	Val	Pro	Val	Asp 395	His	Asp	Leu	Met	Leu 400
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Glu	Ala	Gly	Ser 420	Val	Ala	Ser	Phe	Ile 425	Ala	Ala	Gly	Thr	Ser 430	Tyr	Ala
Thr	His.	Gly 435	Lys	Ala	Met	Ile <sup>.</sup>	Pro 440	Leu	Туг	Ile	Phe	Tyr 445	Ser	Met	Phe
Gly	Ile	Pro	Ala	His	Arg										

148

450

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			atg Met												144
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			gcc Ala												240
			atc Ile												288
		-	aac Asn 100	_		_	_	_	_			_	_		336
		-	atc Ile		-	-					-		-		384
			atc Ile												432
			cag Gln												480
_		-	tac Tyr	-			-	-	-		-		-	_	528
			ttc Phe 180		_	-		_	-	-		-	-		576

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												ggc Gly	720
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												acc Thr	816
												cca Pro	864
												gcg Ala	912
												cag Gln	960
												aag Lys 335	1008
												ctg Leu	1056
Āsp	Gly	Leu	Āla		Arg	Leu	Val	Pro	Ile	Ile	Pro	gat Asp	1104
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Phe Asn Arg Tyr Leu Glu Asn Arg Gly Ile Lys Asp Thr Ser Asp Gln 50 55 60

His Val Trp Ala Phe Leu Gly Asp Gly Glu Met Asp Glu Pro Glu Ser 65 70 75 80

Arg Gly Leu Ile Gln Gln Ala Ala Leu Asn Asn Leu Asp Asn Leu Thr 85 90 95

Phe Val Val Asn Cys Asn Leu Gln Arg Leu Asp Gly Pro Val Arg Gly
100 105 110

Asn Thr Lys Ile Ile Gln Glu Leu Glu Ser Phe Phe Arg Gly Ala Gly
115 120 125

Trp Ser Val Ile Lys Val Val Trp Gly Arg Glu Trp Asp Glu Leu Leu 130 135 140

Glu Lys Asp Gln Asp Gly Ala Leu Val Glu Ile Met Asn Asn Thr Ser 145 150 155 160

Asp Gly Asp Tyr Gln Thr Phe Lys Ala Asn Asp Gly Ala Tyr Val Arg 165 170 175

Glu His Phe Phe Gly Arg Asp Pro Arg Thr Ala Lys Leu Val Glu Asn 180 185 190

Met Thr Asp Glu Glu Ile Trp Lys Leu Pro Arg Gly Gly His Asp Tyr 195 200 205

Arg Lys Val Tyr Ala Ala Tyr Lys Arg Ala Leu Glu Thr Lys Asp Arg 210 220

Pro Thr Val Ile Leu Ala His Thr Ile Lys Gly Tyr Gly Leu Gly His 225 230 235 240

Asn Phe Glu Gly Arg Asn Ala Thr His Gln Met Lys Lys Leu Thr Leu 245 250 255

Asp Asp Leu Lys Leu Phe Arg Asp Lys Gln Gly Ile Pro Ile Thr Asp 260 265 270

Glu Gln Leu Glu Lys Asp Pro Tyr Leu Pro Pro Tyr Tyr His Pro Gly 275 280 285

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Gly Gly Tyr Leu Pro Glu Arg Arg Glu Asn Tyr Asp Pro Ile Gln Val 305 310 315 320
Pro Pro Leu Asp Lys Leu Arg Ser Val Arg Lys Gly Ser Gly Lys Gln 325 330 335
Gln Ile Ala Thr Thr Met Ala Thr Val Arg Thr Phe Lys Glu Leu Met 340 345 350
Arg Asp Lys Gly Leu Ala Asp Arg Leu Val Pro Ile Ile Pro Asp Glu 355 360 365
Ala Arg Thr Phe Gly Leu Asp Ser Trp Phe Pro Thr Leu Lys Ile Tyr 370 375 380
Asn Pro His Gly Gln Asn Tyr Val Pro Val Asp His Asp Leu Met Leu 385 390 395 400
Ser Tyr Arg Glu Ala Pro Glu Gly Gln Ile Leu His Glu Gly Ile Asn 405 410 415
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								ctg Leu								546
								gac Asp								594
								ctc Leu								642
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gtg	act	gta	ctg	acc	tat	tcc	cac	tgg	cgt	ccg	ttg	cag	cga	tgc	agg	1074

Val Thr Val Leu Thr Tyr Ser His Trp Arg Pro Leu Gln Arg Cys Arg 315 320 325

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Asp Val Gly Val Gly Ser Ala Val Thr Met Asp Cys Val Pro Ser 35 40 45

Lys Ser Phe Ile Ala Gly Thr Gly Ile Lys Thr Asp Leu Arg Arg Ala
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Asp Ala Leu Asn Ile Arg Val Lys Asp Leu Ala Lys Ala Gln Ser Glu 85 90 95

Asp Ile Leu Gly Gln Leu Gln Arg Ser Asp Val Arg Met Ile Asn Gly 100 105 110

Val Gly Arg Phe Asp Asp Tyr Asn Thr Lys Gln Thr Thr His Tyr Ile 115 120 125

Lys Val Thr His Ser Asp Gly Ser Glu Glu Thr Val Glu Cys Asp Leu 130 135 140

Val Leu Val Ala Thr Gly Ala Thr Pro Arg Ile Leu Lys Gly Ala Glu 145 150 155 160

Pro Asp Gly Glu Arg Ile Leu Thr Trp Arg Gln Val Tyr Asp Ile Glu 165 170 175

Glu Leu Pro Thr His Leu Ile Val Val Gly Ser Gly Val Thr Gly Ala 180 185 190

Glu Phe Val Ser Ala Phe Ala Glu Leu Gly Val Lys Val Thr Met Val 195 200 205

Ala Ser Arg Asp Arg Ile Leu Pro His Asp Asp Ala Asp Ala Asp 210 215 220

Val Leu Glu Thr Val Leu Ala Glu Arg Gly Val Ser Leu Glu Lys His 225 230 235 240

Ala Arg Val Glu Ser Val Thr Arg Thr Glu Asp Gly Gly Val Cys Val Arg Thr Ala Asp Gly Arg Glu Ile Tyr Gly Ser His Ala Leu Met Thr Val Gly Ser Ile Pro Asn Thr Ala Asp Leu Gly Leu Glu Asn Ile Gly 280 Val Glu Leu Ala Pro Ser Gly His Ile Lys Val Asp Arg Ser Pro Ala Pro Thr Ser Pro Val Cys Thr Gln Gln Val Thr Val Leu Thr Tyr Ser His Trp Arg Pro Leu Gln Arg Cys Arg Ala Val Ser Pro Cys Ile Thr 330 His Ser Val Lys Ala 340 <210> 107 <211> 1518 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (89)..(1495) <223> FRXA02853 <400> 107 aattcagcag taatcattta gacttggaac cgcttaccag tggtttcaac aatgcattca 60 cccagctcac acgtgtggag gtgccttaatg gca aag agg atc gta att atc ggc 115 Met Ala Lys Arg Ile Val Ile Ile Gly 163 ggt gga cet gea gge tat gaa gee gea ete gea gge get aaa tae ggt Gly Gly Pro Ala Gly Tyr Glu Ala Ala Leu Ala Gly Ala Lys Tyr Gly gca gaa gtt acc gtt att gaa gat gtc gga gtt ggc gga tcc gca gtc 211 Ala Glu Val Thr Val Ile Glu Asp Val Gly Val Gly Ser Ala Val 35 acc atg gac tgt gta cct tca aag tcc ttc atc gct ggt acc ggt atc 259 Thr Met Asp Cys Val Pro Ser Lys Ser Phe Ile Ala Gly Thr Gly Ile 307 aaa acc gac ctc cga cgt gct gat gac atg gga ctt aac cgt ggg ctt Lys Thr Asp Leu Arg Arg Ala Asp Asp Met Gly Leu Asn Arg Gly Leu 355 qqa aaa qca cac cta qaa atc qat qca ctq aac atc cqt gtg aag gac Gly Lys Ala His Leu Glu Ile Asp Ala Leu Asn Ile Arg Val Lys Asp 403 ctt qcq aaa qca caq tcc qaa qat atc ttq qqc caq ctq cag cgc tca Leu Ala Lys Ala Gln Ser Glu Asp Ile Leu Gly Gln Leu Gln Arg Ser

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					att Ile 175											643
					ggt Gly											691
					atg Met											739
					gca Ala											787
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100 105 110

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Val Ala Pro Thr Ala Ser Glu Leu Ile Leu Pro Ile Ala Val Ala Val 420 425 430

Lys Leu Phe Cys Arg Arg Asn Ser Gly Leu Ile Ile Gly Gly Val Val

410

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		tac aag ctg Tyr Lys Leu 220			
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					ctt Leu											1363
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					tcc Ser										2371
					cag Gln										2419
					gag Glu										2467
	-		-	Glu	ctg Leu 795	Gln		Asn		Ser					2515
					aac Asn										2563
					tac Tyr										2611
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200

195

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Glu	Arg	Phe	Gly	Glu 245	Gly	Val	Pro	Leu	Lys 250	Pro	Val	Val	Lys	Pro 255	Gly
Ser	Trp	Ile	Gly 260	Gly	Asp	His	Asp	Gly 265	Asn	Pro	Tyr	Val	Thr 270	Ala	Glu
Thr	Val	Glu 275	Tyr	Ser	Thr	His	Arg 280	Ala	Ala	Glu	Thr	Val 285	Leu	Lys	Tyr
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His	Gly	Val	Arg 340	Gly	Arg	Ile	Leu	Ala 345	Thr	Thr	Ala	Glu	Leu 350	Ile	Gly
Glu	Asp	Ala 355	Val	Glu	Gly	Val	Trp 360	Phe	Lys	Val	Phe	Thr 365	Pro	Tyr	Ala
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Ile	Ser	Ala	Ile	Glu 405	Ser	Phe	Gly	Phe	Asn 410	Leu	Tyr	Ala	Leu	Asp 415	Leu
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Arg	Ala	Gln 435	Val	Thr	Ala	Asn	Tyr 440	Arg	Glu	Leu	Ser	Glu 445	Ala	Glu	Lys
Leu	Glu 450	Val	Leu	Leu	Lys	Glu 455	Leu	Arg	Ser	Pro	Arg 460	Pro	Leu	Ile	Pro
His 465	Gly	Ser	Asp	Glu	Tyr 470	Ser	Glu	Val	Thr	Asp 475	Arg	Glu	Leu	Gly	Ile 480
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Ala	Val	Gln	Gly	Ser 645	Val	Arg	Ile	Thr	Glu 650	Gln	Gly	Glu	Ile	Ile 655	Ser
Ala	Lys	Tyr	Gly 660	Asn	Pro	Glu	Thr	Ala 665	Arg	Arg	Asn	Leu	Glu 670	Ala	Leu
Val	Ser	Ala 675	Thr	Leu	Glu	Ala	Ser 680	Leu	Leu	Asp	Val	Ser 685	Glu	Leu	Thr
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Gly 785	Glu	Gln	Ala	Thr	Gln 790	Arg	Ile	Ala	Glu	Leu 795	Gln	Thr	Leu	Asn	Glu 800
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Ser	Lys	Ala	Glu 820	Leu	Arg	Leu	Ala	Lys 825	Leu	Tyr	Ala	Asp	Leu 830	Ile	Pro
Asp	Thr	Glu 835	Val	Ala	Glu	Arg	Val 840	Tyr	Ser	Val	Ile	Arg 845	Glu	Glu	Tyr
Phe	Leu	Thr	Lys	Lys	Met	Phe	Cys	Val	Ile	Thr	Gly	Ser	Asp	Asp	Leu

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Awa The Awa Al-		l Awa Com (	Glu Gly Lys Ala Pro Leu Thr	

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Arg Thr Arg Ala Leu Glu Gly Arg Ser Glu Gly Lys Ala Pro Leu Thr 65 70 75 80

Glu Val Pro Glu Glu Glu Gln Ala His Leu Asp Ala Asp Asp Ser Lys 85 90 95

Glu Arg Arg Asn Ser Leu Asn Arg Leu Leu Phe Pro Lys Pro Thr Glu 100 105 110

Glu Phe Leu Glu His Arg Arg Arg Phe Gly Asn Thr Ser Ala Leu Asp 115 120 125

Asp Arg Glu Phe Phe Tyr Gly Leu Val Glu Gly Arg Glu Thr Leu Ile 130 135 140

Arg Leu Pro Asp Val Arg Thr Pro Leu Leu Val Arg Leu Asp Ala Ile 145 150 155 160

Ser Glu Pro Asp Asp Lys Gly Met Arg Asn Val Val Ala Asn Val Asn
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Gly Gln Ile Arg Pro Met Arg Val Arg Asp Arg Ser Val Glu Ser Val
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Thr Ala Thr Ala Glu Lys Ala Asp Ser Ser Asn Lys Gly His Val Ala 195 200 205

Ala Pro Phe Ala Gly Val Val Thr Val Thr Val Ala Glu Gly Asp Glu 210 215 220

Val Lys Ala Gly Asp Ala Val Ala Ile Ile Glu Ala Met Lys Met Glu 225 230 235 240

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														cgc Arg		307
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					Ala									atc Ile 100		403
														gca Ala		451
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					ctg Leu											643
					gaa Glu											691
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					ttg Leu 235											835
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					cgc Arg											979
ttg Leu	tcc Ser 295	aac Asn	ctg Leu	cgt Arg	gca Ala	cag Gln 300	gcc Ala	acc Thr	gca Ala	ctg Leu	ggc Gly 305	ctt Leu	gcg Ala	gat Asp	cgt Arg	1027
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caa																1083
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Ala	Val	Ala	Lys 20	Leu	Thr	Pro	Glu	Leu 25	Leu	Ser	Val	Glu	Ala 30	Trp	Gly	

Gly Ala Thr Tyr Asp Val Ala Met Arg Phe Leu Phe Glu Asp Pro Trp 35 40 45

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174

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ggt gca gat gct Gly Ala Asp Ala 215	Val Asp (							787
ctc cca agc cat Leu Pro Ser His 230			Ile Val					835
cgt cgc gat acc Arg Arg Asp Thr	ggt ttg a Gly Leu S 250	agc ctc g Ser Leu G	gag gct Glu Ala 255	gtt tct Val Ser	gac ( Asp 1	ctc gag Leu Glu 260	Pro	883
tac tgg gaa gca Tyr Trp Glu Ala 265		Gly Leu T			Glu S			931
cca ggc cca acc Pro Gly Pro Thr 280								979
ttg tcc aac ctg Leu Ser Asn Leu 295	Arg Ala		_				_	1027
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ctt ggc ggt ctc ttg ttg aaa gga ata att act cta gtg tcg act cac 163
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gcc Ala	acg Thr 55	gta Val	gct Ala	att Ile	tac Tyr	ccc Pro 60	cgt Arg	gaa Glu	gat Asp	cgg Arg	gga Gly 65	tca Ser	ttc Phe	cac His	cgc Arg	307
				gaa Glu												355
				gac Asp 90												403
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				gag Glu												499
				ctt												547
	Pro 135	Glu	Val	Leu	Asp	Leu 140	Thr	Gly	Asp	гуѕ	145	Arg	Ala	Val	Thr	
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gcc Ala 150 aaa Lys	gcg Ala aac Asn	aag Lys atc Ile	aag Lys gat Asp	gct Ala gag Glu	ggt Gly 155 atc Ile	ctg Leu gtt Val	cca Pro aaa Lys	gtt Val agc Ser	ttg Leu gct Ala 175	gcg Ala 160 gaa Glu	gaa Glu ggc Gly	tcc Ser cag Gln	acc Thr act Thr	ccg Pro tac Tyr 180	agc Ser 165 ccc Pro	
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gcc Ala 150 aaa Lys atc Ile gtt Val gaa Glu	gcg Ala aac Asn ttt Phe gct Ala gct Ala 215 att	aag Lys atc Ile gtg Val tca Ser 200 gaa Glu	aag Lys gat Asp aag Lys 185 cct Pro	gct Ala gag Glu 170 gca Ala gat Asp	ggt Gly 155 atc Ile gtt Val gag Glu ttc Phe	ctg Leu gtt Val gcc Ala ctt Leu ggc Gly 220 att	cca Pro aaa Lys ggt Gly cgc Arg 205 gat Asp	gtt Val agc Ser ggt Gly 190 aaa Lys ggc Gly	ttg Leu gct Ala 175 ggc Gly tta Leu gcg Ala	gcg Ala 160 gaa Glu gga Gly gca Ala gta Val	gaa Glu ggc Gly cgc Arg aca Thr tat Tyr 225	tcc Ser cag Gln ggt Gly gaa Glu 210 gtc Val	acc Thr act Thr atg Met 195 gca Ala gaa Glu	ccg Pro tac Tyr 180 cgt Arg tct Ser cgt Arg	agc Ser 165 ccc Pro ttt Phe cgt Arg	643 691 739

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ggt tac cag ggc Gly Tyr Gln Gly 295	gcg gga acc Ala Gly Thr 300	gtg gaa Val Glu	ttc ttg gtc ga Phe Leu Val As 305	at gaa aag sp Glu Lys	ggc 1027 Gly
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atc aag acc cac Ile Lys Thr His 360	ggt gca gca £ly Ala Ala	ctg cag Leu Gln 365	Cys Arg Ile Th	cc acg gaa nr Thr Glu 70	gat 1219 Asp
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ggc gaa atc acc Gly Glu Ile Thr		Asp Ser			
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ttc att gcc gat Phe Ile Ala Asp 470					
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His	Leu	Asp 275	Pro	Glu	Leu	Arg	Asp 280	Arg	Ile	Cys	Ala	Asp 285	Ala	Val	Lys
Phe	Cys 290	Arg	Ser	Ile	Gly	Tyr 295	Gln	Gly	Ala	Gly	Thr 300	Val	Glu	Phe	Leu
Val 305	Asp	Glu	Lys	Gly	Asn 310	His	Val	Phe	Ile	Glu 315	Met	Asn	Pro	Arg	Ile 320
Gln	Val	Glu	His	Thr 325	Val	Thr	Glu	Glu	Val 330	Thr	Glu	Val	Asp	Leu 335	Val
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Arg 465	Ile	Ala	Thr	Gly	Phe 470	Ile	Ala	Asp	His	Pro 475	His	Leu	Leu	Gln	Ala 480
Pro	Pro	Ala	Asp	Asp 485	Glu	Gln	Gly	Arg	Ile 490	Leu	Asp	Tyr	Leu	Ala 495	Asp
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		gcg Ala													720
		atc Ile													768
		gtc Val													816
		gca Ala 275													864
		ggt Gly	_	-	-	_	_	-			_	-	-		912
		ttc Phe	_		_								_		960
		gct Ala													1008
		gca Ala			•		_	-			_		_	_	 1056
		ttt Phe 355													1104
		gtg Val	Ser	Gly	Val		Thr	Asn	Ile	Gly	Phe	Leu			1152
		gaa Glu													1200
		cac His													1248
		atc Ile													1296
		cgt Arg 435													1344

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Ala Arg Glu Cys Ala Glu Asn Gly Ile Thr Phe Ile Gly Pro Thr Pro 50 55 60

Glu Val Leu Asp Leu Thr Gly Asp Lys Ser Arg Ala Val Thr Ala Ala 65 70 75 80

Lys Lys Ala Gly Leu Pro Val Leu Ala Glu Ser Thr Pro Ser Lys Asn 85 90 95

Ile Asp Glu Ile Val Lys Ser Ala Glu Gly Gln Thr Tyr Pro Ile Phe 100 105 110

Val Lys Ala Val Ala Gly Gly Gly Gly Arg Gly Met Arg Phe Val Ala 115 120 125

Ser Pro Asp Glu Leu Arg Lys Leu Ala Thr Glu Ala Ser Arg Glu Ala 130 135 140

Glu Ala Ala Phe Gly Asp Gly Ala Val Tyr Val Glu Arg Ala Val Ile 145 150 155 160

Asn Pro Gln His Ile Glu Val Gln Ile Leu Gly Asp His Thr Gly Glu 165 170 175

Val Val His Leu Tyr Glu Arg Asp Cys Ser Leu Gln Arg Arg His Gln 180 185 190

Lys Val Val Glu Ile Ala Pro Ala Gln His Leu Asp Pro Glu Leu Arg 195 200 205

Asp Arg Ile Cys Ala Asp Ala Val Lys Phe Cys Arg Ser Ile Gly Tyr 210 215 220

Gln Gly Ala Gly Thr Val Glu Phe Leu Val Asp Glu Lys Gly Asn His 225 230 235 240

Val Phe Ile Glu Met Asn Pro Arg Ile Gln Val Glu His Thr Val Thr  $245 \hspace{1.5cm} 250 \hspace{1.5cm} 255$ 

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Thr	His 290	Gly	Ala	Ala	Leu	Gln 295	Cys	Arg	Ile	Thr	Thr 300	Glu	Asp	Pro	Asn	
Asn 305	Gly	Phe	Arg	Pro	Asp 310	Thr	Gly	Thr	Ile	Thr 315	Ala	Tyr	Arg	Ser	Pro 320	
Gly	Gly	Ala	Gly	Val 325	Arg	Leu	Asp	Gly	Ala 330	Ala	Gln	Leu	Gly	Gly 335	Glu	
Ile	Thr	Ala	His 340	Phe	Asp	Ser	Met	Leu 345	Val	Lys	Met	Thr	Cys 350	Arg	Gly	
Ser	Asp	Phe 355	Glu	Thr	Ala	Val	Ala 360	Arg	Ala	Gln	Arg	Ala 365	Leu	Ala	Glu	
Phe	Thr 370	Val	Ser	Gly	Val	Ala 375	Thr	Asn	Ile	Gly	Phe 380	Leu	Arg	Ala	Leu	
Leu 385	Arg	Glu	Glu	Asp	Phe 390	Thr	Ser	Lys	Arg	Ile 395	Ala	Thr	Gly	Phe	Ile 400	
Ala	Asp	His	Pro	His 405	Leu	Leu	Gln	Ala	Pro 410	Pro	Ala	Asp	Asp	Glu 415	Gln	
Gly	Arg	Ile	Leu 420	Asp	Tyr	Leu	Ala	Asp 425	Val	Thr	Val	Asn	Lys 430	Pro	His	
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	gat Asp															259
	atc Ile 55															307
	acc Thr															355
	atc Ile															403
	ttt Phe															451
-	cac His	-	-	-	_	_	-				-	-				499
	ttc Phe 135															547
	gtg Val															595
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	gga Gly 215															787
	agc Ser															835
	aac Asn															883
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ttc cca gga Phe Pro Gly						: Ile
acc ccc gag Thr Pro Glu						
ctg agg acc Leu Arg Thr 360			a Ser Cys	Leu Pro P		
gcg ttg ccc Ala Leu Pro 375					er Pro Ly	
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35	-	4	)	ı	45	
Ala Gln Val 50	Cys Glu	Ala Ile Ly: 55	s Glu Asp	Pro Glu Va 60	al Ala Arq	g Thr
His Thr Gly						

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Glu	Gly	Lys	Ala 100	Gln	Leu	Phe	Ser	Ser 105	Phe	Ala	Gly	Leu	Lys 110	Ala	Ile
Pro	Ile	Val 115	Leu	Asp	Val	His	Asp 120	Val	Asp	Ala	Leu	Val 125	Glu	Thr	Ile
Ala	Ala 130	Ile	Ala	Pro	Ser	Phe 135	Gly	Ala	Ile	Asn	Leu 140	Glu	Asp	Ile	Ser
Ala 145	Pro	Arg	Cys	Phe	Glu 150	Val	Glu	Arg	Arg	Leu 155	Ile	Glu	Arg	Leu	Asp 160
Ile	Pro	Val	Met	His 165	Asp	Asp	Gln	His	Gly 170	Thr	Ala	Val	Val	Ile 175	Leu
Ala	Ala	Leu	Arg 180	Asn	Ser	Leu	Lys	Leu 185	Leu	Asp	Arg	Lys	Ile 190	Glu	Asp
Leu	Lys	Ile 195	Val	Ile	Ser	Gly	Ala 200	Gly	Ala	Ala	Gly	Val 205	Ala	Ala	Val
Asp	Met 210	Leu	Thr	Asn	Ala	Gly 215	Ala	Thr	Asp	Ile	Val 220	Val	Leu	Asp	Ser
Arg 225	Gly	Ile	Ile	His	Asp 230	Ser	Arg	Glu	Asp	Leu 235	Ser	Pro	Val	Lys	Ala 240
Ala	Leu	Ala	Glu	Lys 245	Thr	Asn	Pro	Arg	Gly 250	Ile	Ser	Gly	Gly	Ile 255	Asn
Glu	Ala	Phe	Thr 260	Gly	Ala	Asp	Leu	Phe 265	Ile	Gly	Val	Ser	Gly 270	Gly	Asn
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Tyr 305	Gly	Ala	Ile	Val	Ala 310	Thr	Gly	Arg	Ser	Asp 315	Leu	Pro	Asn	Gln	Ile 320
Asn	Asn	Val	Leu	Ala 325	Phe	Pro	Gly	Ile	Phe 330	Ala	Gly	Ala	Leu	Ala 335	Ala
Lys	Ala	Lys	Lys 340	Ile	Thr	Pro	Glu	Met 345	Lys	Leu	Ala	Ala	Gln 350	Arg	Gln
Ser	Gln	Thr 355	Ser	Gln	Leu	Arg	Thr 360	Ser	Arg	Ser	Ala	Ala 365	Ser	Cys	Leu
Pro	Pro 370	Trp	Ile	Pro	Ala	Leu 375	Pro	Gln	Gln	Ser	Arg 380	Gln	Leu	Ser	Arg
Pro 385	Ser	Pro	Lys	Arg	Lys 390	Thr	Leu	Lys	Asn	Leu 395	Leu	Ile	Asp	Ala	Ser 400
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	gac Asp															931
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Pro	Leu	Arg 35	Asp	Met	Arg	Asp	Leu 40	Ser	Leu	Ala	Tyr	Thr 45	Pro	Gly	Val	
Ala	Gln	Val	Cys	Glu	Ala	Ile	Lys	Glu	Asp	Pro	Glu	Val	Ala	Arg	Thr	

50 55 60

His Thr Gly Ile Gly Asn Thr Val Ala Val Ile Ser Asp Gly Thr Ala 65 70 75 80

Val Leu Gly Leu Gly Asp Ile Gly Pro Gln Ala Ser Leu Pro Val Met 85 90 95

Glu Gly Lys Ala Gln Leu Phe Ser Ser Phe Ala Gly Leu Lys Ala Ile 100 \$105\$

Pro Ile Val Leu Asp Val His Asp Val Asp Ala Leu Val Glu Thr Ile 115 120 125

Ala Ala Ile Ala Pro Ser Phe Gly Ala Ile Asn Leu Glu Asp Ile Ser 130 135 140

Ala Pro Arg Cys Phe Glu Val Glu Arg Arg Leu Ile Glu Arg Leu Asp 145 150 155 160

Ile Pro Val Met His Asp Asp Gln His Gly Thr Ala Val Val Ile Leu 165 170 175

Ala Ala Leu Arg Asn Ser Leu Lys Leu Leu Asp Arg Lys Ile Glu Asp 180 185 190

Leu Lys Ile Val Ile Ser Gly Ala Gly Ala Ala Gly Val Ala Ala Val 195 200 205

Asp Met Leu Thr Asn Ala Gly Ala Thr Asp Ile Val Val Leu Asp Ser 210 215 220

Arg Gly Ile Ile His Asp Ser Arg Glu Asp Leu Ser Pro Val Lys Ala 225 230 235 240

Ala Leu Ala Glu Lys Thr Asn Pro Arg Gly Ile Ser Gly Gly Ile Asn 245 250 255

Glu Ala Phe Thr Gly Ala Asp Leu Phe Ile Gly Val Ser Gly Gly Asn 260 265 270

Ile Gly Glu Asp Ala Leu Lys Leu Met Ala Pro Glu Pro Ile Leu Phe 275 280 285

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ggc tcc act tcc tac ggc Gly Ser Thr Ser Tyr Gl 230 23	/ Ile Gly Met			5
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190

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cct ggc gat Pro Gly Asp		ly Gly Ala		787
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ggt gag cgt Gly Glu Arg 250				883
ctt att gtg Leu Ile Val 265	Ala Lys As			931
ccg cag gcg Pro Gln Ala				979
cgt att tat Arg Ile Tyr		lu Asp Val		1027
agc aag gcg Ser Lys Ala 315	Ala Lys A			1075
tat gag atg Tyr Glu Met 330				1123
 acc ttt gtt Thr Phe Val 345	Asp Gln A	_	 	1171
ggc aag tca Gly Lys Ser		sp Ile Gly		1219
ttg gcg gat Leu Ala Asp		lu Gly Thr		1267
ggg ccg gtg Gly Pro Val 395				1315
gat aag gca Asp Lys Ala 410			J . J	1363
tcg tcg gaa Ser Ser Glu 425	Thr Gly As			1411

Ala Gly Gly 440	atc ggt Ile Gly	att aat Ile Asn	gat ggc Asp Gly 445	tac gcc Tyr Ala	gcg acg Ala Thr 450	tgg gc Trp Al	g agc a Ser	1459
gtg tcc acg Val Ser Thr 455								1507
cat ggt gcg His Gly Ala 470	gag gga Glu Gly	att aca Ile Thr 475	aaa tat Lys Tyr	gcg gag Ala Glu 480	atc cga Ile Arg	aac at Asn Il	c gcg e Ala 485	1555
gag cag cgc Glu Gln Arg							g Lys	1603
gtg tac tca Val Tyr Ser	gac acc Asp Thr 505	gtg gcc Val Ala	aca gcg Thr Ala 510	Leu Lys	ctg ggc Leu Gly	aaa at Lys Il 515	c ttt e Phe	1651
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Val Ile Ala 35	Pro Phe	Thr Gly	Glu Thr	Leu Gly	Phe Gly 45		p Gly	
			40		45			
35 Asp Glu Gln	Asp Val	Glu His 55	40 Ala Phe	Ala Leu	Ser Arg 60 Lys Lys	Ala Al	a Gln	
Asp Glu Gln 50  Lys Lys Trp	Asp Val Val His	Glu His 55 Thr Thr 70	40 Ala Phe Ala Val	Ala Leu Glu Arg 75	Ser Arg 60 Lys Lys	Ala Al Ile Ph	a Gln e Leu 80	
Asp Glu Gln 50 Lys Lys Trp 65	Asp Val Val His Asp Leu 85	Glu His 55 Thr Thr 70 Val Leu	40 Ala Phe Ala Val	Ala Leu Glu Arg 75 Arg Glu 90 Ala Ser	Ser Arg 60 Lys Lys Leu Leu	Ala Al Ile Ph Met As	a Gln e Leu 80 p Ile 5	
Asp Glu Gln 50  Lys Lys Trp 65  Lys Val His	Asp Val Val His Asp Leu 85 Glu Thr	Glu His 55 Thr Thr 70 Val Leu Gly Lys	Ala Phe Ala Val Lys Asr Asn Arg	Ala Leu Glu Arg 75 Arg Glu 90 Ala Ser	Ser Arg 60 Lys Lys Leu Leu Ala Ala	Ala Al Ile Ph Met As 9 Asp Gl	a Gln e Leu 80 p Ile 5	
Asp Glu Gln 50  Lys Lys Trp 65  Lys Val His  Val Gln Leu  Leu Asp Val	Asp Val Val His Asp Leu 85 Glu Thr 100 Ala Ile	Glu His 55 Thr Thr 70 Val Leu Gly Lys Thr Thr	Ala Phe Ala Val Lys Asr Asn Arg 105 Arg Phe	Ala Leu Glu Arg 75 Arg Glu 90 Ala Ser	Ser Arg 60 Lys Lys Leu Leu Ala Ala Asn Asn 125	Ala Al Ile Ph Met As 9 Asp Gl 110 Ala Gl	a Gln e Leu 80 p Ile 5	

Trp	Asn	Tyr	Pro	Leu 165	Thr	Leu	Gly	Val	Ser 170	Asp	Ala	Val	Pro	Ala 175	Leu
Leu	Ala	Gly	Asn 180	Ala	Val	Val	Ala	Lys 185	Pro	Asp	Leu	Ala	Thr 190	Pro	Phe
Ser	Cys	Leu 195	Ile	Met	Val	His	Leu 200	Leu	Ile	Glu	Ala	Gly 205	Leu	Pro	Arg
Asp	Leu 210	Met	Gln	Val	Val	Thr 215	Gly	Pro	Gly	Asp	Ile 220	Val	Gly	Gly	Ala
Ile 225	Ala	Ala	Gln	Cys	Asp 230	Phe	Leu	Met	Phe	Thr 235	Gly	Ser	Thr	Ala	Thr 240
Gly	Arg	Ile	Leu	Gly 245	Arg	Thr	Met	Gly	Glu 250	Arg	Leu	Val	Gly	Phe 255	Ser
Ala	Glu	Leu	Gly 260	Gly	Lys	Asn	Pro	Leu 265	Ile	Val	Ala	Lys	Asp 270	Ala	Asp
Leu	Asp	Lys 275	Val	Glu	Ala	Glu	Leu 280	Pro	Gln	Ala	Cys	Phe 285	Ser	Asn	Ser
Gly	Gln 290	Leu	Cys	Val	Ser	Thr 295	Glu	Arg	Ile	Tyr	Val 300	Glu	Glu	Asp	Val
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Ile	Gly	Ala	Gly	Phe 325	Glu	Trp	Lys	Tyr	Glu 330	Met	Gly	Ser	Leu	11e 335	Asn
His	Ala	Gln	Leu 340	Asp	Arg	Val	Ser	Thr 345	Phe	Val	Asp	Gln	Ala 350	Lys	Ala
Ala	Gly	Ala 355	Thr	Val	Leu	Суѕ	Gly 360	Gly	Lys	Ser	Arg	Pro 365	Asp	Ile	Gly
Pro	Phe 370	Phe	Tyr	Glu	Pro	Thr 375	Val	Leu	Ala	Asp	Val 380	Pro	Glu	Gly	Thr
Pro 385	Leu	Leu	Thr	Glu	Glu 390	Val	Phe	Gly	Pro	Val 395	Val	Phe	Ile	Glu	Lys 400
Val	Ala	Thr	Leu	Glu 405	Glu	Ala	Val	Asp	Lys 410	Ala	Asn	Gly	Thr	Pro 415	Tyr
Gly	Leu	Asn	Ala 420	Ser	Val	Phe	Gly	Ser 425	Ser	Glu	Thr	Gly	Asn 430	Leu	Val
Ala	Gly	Gln 435	Leu	Glu	Ala	Gly	Gly 440	Ile	Gly	Ile	Asn	Asp 445	Gly	Tyr	Ala
Ala	Thr 450	Trp	Ala	Ser	Val	Ser 455	Thr	Pro	Leu	Gly	Gly 460	Met	Lys	Gln	Ser
Gly 465	Leu	Gly	His	Arg	His 470	Gly	Ala	Glu	Gly	Ile 475	Thr	Lys	Tyr	Ala	Glu 480
Ile	Arg	Asn	Ile	Ala	Glu	Gln	Arg	Trp	Met	Ser	Met	Arg	Gly	Pro	Ala

490

485

495

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gtc gga Val Gly 150														595
tcg gtt Ser Va														643
gcg agt Ala Se														691
gag gag Glu Glu														739
gga to Gly Ser 21	Glu													787
att to Ile Se 230														835
gca att Ala Ile														883
aac gco Asn Ala														931
cag gct Gln Ala														979
tca ato Ser Ile 29	e Asn	-	_		-	-	-	-	-		-	_		1027
gag aad Glu Lys 310	-	Val	Glu	_	Val	Lys	Asn	Ile	Pro	Thr		-	-	1075
gca gaa Ala Gli														1123
ggt tte														1171
cag gti Gln Va														1219
tct gas Ser As <sub>1</sub> 37	Val													1267

cct	ctc	atc	agc	gtg	ctg	aag	gcc	gat	gat	gag	gca	cac	gca	gca	gag	1315
Pro 390	Leu	Ile	Ser	Val	Leu 395	Lys	Ala	Asp	Asp	Glu 400	Ala	His	Ala	Ala	Glu 405	
	gcc Ala															1363
	att Ile															1411
	cac His															1459
	ggt Gly 455		-						_				-			1507
	gag Glu					-	_			-		-	_	_		1552
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150 155 145 Asn Leu Ser Ile Arg Ser Val Ala Pro Ala Leu Ala Val Gly Asn Ala 170 Val Val Ile Lys Pro Ala Ser Asp Thr Pro Val Thr Gly Gly Val Ile 180 185 Pro Ala Arg Ile Phe Glu Glu Ala Gly Val Pro Ala Gly Val Ile Ser 200 Thr Val Ala Gly Ala Gly Ser Glu Ile Gly Asp His Phe Val Thr His 215 Ala Val Pro Lys Leu Ile Ser Phe Thr Gly Ser Thr Pro Val Gly Arg 230 Arg Val Gly Glu Leu Ala Ile Asn Gly Gly Pro Met Lys Thr Val Ala Leu Glu Leu Gly Gly Asn Ala Pro Phe Val Val Leu Ala Asp Ala Asp Ile Asp Ala Ala Ala Gln Ala Ala Val Gly Ala Phe Leu His Gln Gly Gln Ile Cys Met Ser Ile Asn Arg Val Ile Val Asp Ala Ala Val His Asp Glu Phe Leu Glu Lys Phe Val Glu Ala Val Lys Asn Ile Pro Thr Gly Asp Pro Ser Ala Glu Gly Thr Leu Val Gly Pro Val Ile Asn 330 Asp Ser Gln Leu Ser Gly Leu Lys Glu Lys Ile Glu Leu Ala Lys Lys Glu Gly Ala Thr Val Gln Val Glu Gly Pro Ile Glu Gly Arg Leu Val 360 His Pro His Val Phe Ser Asp Val Thr Ser Asp Met Glu Ile Ala Arg Glu Glu Ile Phe Gly Pro Leu Ile Ser Val Leu Lys Ala Asp Asp Glu Ala His Ala Ala Glu Leu Ala Asn Ala Ser Asp Phe Gly Leu Ser Ala 405 Ala Val Trp Ser Lys Asp Ile Asp Arg Ala Ala Gln Phe Ala Leu Gln 425 Ile Asp Ser Gly Met Val His Ile Asn Asp Leu Thr Val Asn Asp Glu 435 440 Pro His Val Met Phe Gly Gly Ser Lys Asn Ser Gly Leu Gly Arg Phe Asn Gly Asp Trp Ala Ile Glu Glu Phe Thr Thr Asp Arg Trp Ile Gly 470 475

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			g atg cgt gtg a Met Arg Val 175		
			g ctt cca gaa a Leu Pro Glu )		
	Asp Val Pro		c ttg cag gat e Leu Gln Asp		
			gcg ttg gct Ala Leu Ala 225	Arg Ala Ile	
-		Asn Pro Met	g atc att gct Ile Ile Ala 240		
			g cag gca ctt 1 Gln Ala Leu 255		
			t ggt ggt ggc a Gly Gly Gly )		
	Gln Asn Le		g ggt gcc acc l Gly Ala Thr		
			gtg atc atc Val Ile Ile 305	Gly Ile Gly	
		Thr Ala Se	c cgc act gcg r Arg Thr Ala 320		
	Phe Ile Ası	_	gct tcc ttc L Ala Ser Phe 335		Lys
			a gat gca cgc a Asp Ala Arg )		
	a Glu Ala Lei		e acc gtg gca e Thr Val Ala		
			g tgg gac gca a Trp Asp Ala 385	Glu Val Asp	
		Leu Ala Leu	g cct gga cag 1 Pro Gly Gln 400		

ggc gcg g Gly Ala V													1363
gct gga t Ala Gly S													1411
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atc gcg q Ile Ala ( 455													1507
gac cgc o Asp Arg A 470			Met										1555
aac act o													1603
gtg ctc a													1651
act gtc of Thr Val (													1699
gag gcg a Glu Ala I 535													1747
atg aat o Met Asn A 550	-	-	Gly	-	-	_		_	-	-		_	1795
gcg aat o					Ala		Met						1843
tcg gag a Ser Glu 1													1891
gca cca ( Ala Pro /													1939
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<213> Corynebacterium glutamicum

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Pro Gly Met Phe Gly Ile Phe Gly His Gly Asn Val Ala Gly Ile Gly
35 40 45

Gln Ala Leu Lys Gln Tyr Asn Val Glu Gln Pro Glu Leu Met Pro Tyr 50 55 60

Tyr Gln Ala Arg Asn Glu Gln Ala Met Val His Gln Ser Val Gly Tyr
65 70 75 80

Ala Arg Met His Arg Arg Gly Thr Tyr Ala Ser Ala Ala Ser Val 85 90 95

Gly Pro Gly Ala Thr Asn Leu Leu Thr Gly Ala Ala Leu Ala Thr Thr 100 105 110

Asn Arg Leu Pro Ala Leu Leu Pro Ser Asp Thr Phe Ala Thr Arg 115 120 125

Val Ala Asp Pro Val Leu Gln Gln Leu Glu Gln Pro Trp Asp Ile Gly 130 135 140

Leu Thr Val Asn Asp Ala Phe Arg Pro Val Ser Lys Phe Phe Asp Arg 145 150 155 160

Val Gln Arg Pro Glu Gln Leu Phe Ser Ile Ala Leu Ala Ala Met Arg 165 170 175

Val Leu Thr Asp Pro Ala Glu Thr Gly Ala Val Thr Ile Ala Leu Pro 180 185 190

Glu Asp Val Gln Ala Glu Met Leu Asp Val Pro Val Glu Phe Leu Gln 195 200 205

Asp Arg Glu Trp His Ile Arg Arg Pro Arg Pro Glu Arg Ala Ala Leu 210 215 220

Ala Arg Ala Ile Glu Val Ile Lys Asn Ala Lys Asn Pro Met Ile Ile 225 230 235 240

Ala Gly Gly Gly Val Leu Tyr Ser Asp Ala Glu Thr Gln Leu Gln Ala 245 250 255

Leu Val Glu Gln Thr Gly Ile Pro Val Gly Thr Ser Gln Ala Gly Gly 260 265 270

Gly Val Leu Ala Trp Asp His Ala Gln Asn Leu Gly Gly Val Gly Ala 275 280 285

Thr Gly Thr Leu Ala Ala Asn Arg Ile Ala Gly Asp Ala Asp Val Ile

290 295 300

Ile Gly Ile Gly Thr Arg Tyr Ser Asp Phe Thr Thr Ala Ser Arg Thr 305 310 315 320

Ala Phe Gln Asn Pro Asp Val Thr Phe Ile Asn Ile Asn Val Ala Ser 325 330 335

Phe Asp Ala Tyr Lys His Gly Thr Gln Leu Pro Val Ile Ala Asp Ala 340 345 350

Arg Glu Ala Ile Val Glu Leu Ala Glu Ala Leu Gln Gly Phe Thr Val 355 360 365

Ala Glu Asp Tyr Ala Gln Arg Ile Ala Lys Glu Lys Ala Ala Trp Asp 370 380

Ala Glu Val Asp Lys Ser Phe Ala Pro Ser Gly Leu Ala Leu Pro Gly 385 390 395 400

Gln Pro Glu Ile Ile Gly Ala Val Gln Ala Ser Thr Ser Glu Lys Asp 405 410 415

Val Ile Val Gln Ala Ala Gly Ser Leu Pro Gly Asp Leu His Lys Leu 420 425 430

Trp Arg Val Arg Asp Ala Leu Gly Tyr His Val Glu Tyr Ala Phe Ser 435 440 445

Cys Met Gly Tyr Glu Ile Ala Gly Gly Ile Gly Ala Lys Arg Gly Leu 450 455 460

Asp Ala Ala Gly Asp Asp Asp Val Val Ile Met Val Gly Asp Gly 465 470 475 480

Ser Tyr Leu Met Leu Asn Thr Glu Leu Val Thr Ala Val Ala Glu Gly 485 490 495

Ile Lys Val Ile Val Val Leu Ile Gln Asn His Gly Tyr Ala Ser Ile 500 505 510

Gly His Leu Ser Glu Thr Val Gly Ser Gln Arg Phe Gly Thr Trp Tyr 515 520 525

Arg Glu Tyr Asp Ala Glu Ala Lys Asn Phe Gln Gly Glu Gln Ile Leu 530 535 540

Pro Val Asp Leu Ala Met Asn Ala Arg Ser Tyr Gly Met Asp Val Ile 545 550 555 560

Glu Val Glu Pro Ser Ala Asn Ala Ile Glu Asp Leu Lys Ala Ala Met 565 570 575

Ala Thr Ala Lys Ala Ser Glu Lys Ser Thr Phe Ile His Ile Asn Ser 580 585 590

Asp Pro Leu Ile Tyr Ala Pro Asp Gly Ala Gly Trp Trp Asp Val Pro 595 600

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Tyr Leu Lys Asn Gln Ala Leu Gln Arg Pro Leu Leu Gly

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gct Ala																672
acc Thr 225	gcc Ala	gca Ala	gtt Val	gaa Glu	ggt Gly 230	ttc Phe	ccc Pro	att Ile	aag Lys	atc Ile 235	gca Ala	cta Leu	atc Ile	aac Asn	aac Asn 240	720
gga Gly																768
gga Gly																816
gac Asp																864
Thr					gta Val											912
					gtc Val 310											960
_			_		tct Ser	_						_		-		1008
gca Ala			_	-	cca Pro			-		-	-		-	_	_	1056
					cac His											1104
Glu	-		-	_	taaq	ggaga	aga d	ccaa	agato	gg ct	a					1142
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547

gac att cca cag gca ttg gct gag gca ttc cac ctc gcg att act ggt

Asp Ile Pro Gln Ala Leu Ala Glu Ala Phe His Leu Ala Ile Thr Gly

140

135

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gaa f Glu 1																643
cca (																691
atc (	ggt Gly	gag Glu 200	gcc Ala	aag Lys	aag Lys	ccc Pro	gtc Val 205	ctt Leu	tac Tyr	gtt Val	ggt Gly	ggt Gly 210	ggc Gly	gta Val	atc Ile	739
aag ( Lys :	gct Ala 215	gac Asp	gca Ala	cac His	gaa Glu	gag Glu 220	ctt Leu	cgt Arg	gcg Ala	ttc Phe	gct Ala 225	gag Glu	tac Tyr	acc Thr	ggc Gly	787
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Met '	Thr	Gly		5				_	10					15		
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gcc ggt ctg Ala Gly Leu 230					n Ser A				35
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Ile Gln Gly 145	Phe Pro	Ile Leu 150	Ser Ala	Tyr Se		Thr Lys	Phe	Ala 160	
Val Arg Gly	Leu Thr 165	Gln Ala	Ala Ala	Gln Gl 170	u Leu A	Ala Pro	Lys 175	Gly	
His Thr Val	Asn Ala 180	Tyr Ala	Pro Gly 185		l Gly T	Thr Gly 190	Met	Trp	

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				gaa Glu												355
				ttt Phe 90												403
				gtg Val												451
Tyr	Ala	Gly 120	Ğİy	tgg Trp	Ala	Gln	Asn 125	Ile	Val	Val	Pro	Ala 130	Glu	Ala	Leu	499
				gat Asp												547
Cys 150	Āla	ĞÎy	Val	aca Thr	Thr 155	Phe	Asn	Ala	Leu	Arg 160	Asn	Leu	Lys	Leu	Asp 165	595
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		_	-	ttg Leu		_						_				835
				acc Thr 250												883
				gga Gly												931
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Ala Leu Phe Glu Leu Gly Gly Ala Asp Leu Ile Leu Ser Thr Ala Ser 230 Thr Thr Glu Pro Leu Ser Glu Leu Ser Thr Gly Leu Ser Ile Gly Gly Gln Leu Thr Ile Ile Gly Val Asp Gly Gly Asp Ile Thr Val Ser Ala Ala Gln Leu Met Met Asn Arg Gln Ile Ile Thr Gly His Leu Thr Gly Ser Ala Asn Asp Thr Glu Gln Thr Met Lys Phe Ala His Leu His Gly 295 Val Lys Pro Leu Ile Glu Arg Met Pro Leu Asp Gln Ala Asn Glu Ala 315 Ile Ala Arg Ile Ser Ala Gly Lys Pro Arg Phe Arg Ile Val Leu Glu 330 Pro Asn Ser <210> 243 <211> 1665 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(1642) <223> RXA02737 <400> 243 agcacqctgc atcagtaacg gcgacatgaa atcgaattag ttcgatctta tgtggccgtt 60 acacatettt cattaaagaa aggategtga cactaceate gtg age aca aac aeg Val Ser Thr Asn Thr acc ccc tcc agc tgg aca aac cca ctg cgc gac ccg cag gat aaa cga 163 Thr Pro Ser Ser Trp Thr Asn Pro Leu Arg Asp Pro Gln Asp Lys Arg ctc ccc cgc atc gct ggc cct tcc ggc atg gtg atc ttc ggt gtc act 211 Leu Pro Arg Ile Ala Gly Pro Ser Gly Met Val Ile Phe Gly Val Thr 259 gge gae ttg get ega aag aag etg ete eee gee att tat gat eta gea Gly Asp Leu Ala Arg Lys Lys Leu Leu Pro Ala Ile Tyr Asp Leu Ala aac cgc gga ttg ctg ccc cca gga ttc tcg ttg gta ggt tac ggc cgc 307 Asn Arg Gly Leu Leu Pro Pro Gly Phe Ser Leu Val Gly Tyr Gly Arg 355 cgc gaa tgg tcc aaa gaa gac ttt gaa aaa tac gta cgc gat gcc gca Arg Glu Trp Ser Lys Glu Asp Phe Glu Lys Tyr Val Arg Asp Ala Ala

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				ttt Phe										451
				gca Ala										499
	-			gct Ala		_					-			547
				cag Gln 155										595
				cgc Arg										643
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				cgc Arg									gtt <sup>·</sup> Val	739
				ctg Leu										787
				gtt Val 235										835
				cgt Arg										883
				aac Asn										931
_	-	-		tct Ser			-	_	-	_	-	-	_	979
				gcg Ala										1027
				tac Tyr 315										1075

aag gga ctt d Lys Gly Leu <i>l</i>					r Glu
act ttt gcg ( Thr Phe Ala A					
gtg ccg ttc t Val Pro Phe 3					
gag att gcc of Glu Ile Ala v 375	gtg gtg ttt Val Val Phe	aaa gac gca Lys Asp Ala 380	cca cac cag Pro His Gln 385	cct ttc gad Pro Phe As	e gge 1267 e Gly
gac atg act of Asp Met Thr N					
cct gat gaa ( Pro Asp Glu (					y Ser
gcc atg gaa ( Ala Met Glu )					
ttc act gaa ( Phe Thr Glu ( 440					
ctg tta gat ( Leu Leu Asp ( 455					
agc tgg aag a Ser Trp Lys : 470					
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345 350 Arg Arg Trp Ala Gly Val Pro Phe Tyr Leu Arg Thr Gly Lys Arg Leu 360 Gly Arg Arg Val Thr Glu Ile Ala Val Val Phe Lys Asp Ala Pro His 375 380 Gln Pro Phe Asp Gly Asp Met Thr Val Ser Leu Gly Gln Asn Ala Ile 390 Val Ile Arg Val Gln Pro Asp Glu Gly Val Leu Ile Arg Phe Gly Ser Lys Val Pro Gly Ser Ala Met Glu Val Arg Asp Val Asn Met Asp Phe 425 Ser Tyr Ser Glu Ser Phe Thr Glu Glu Ser Pro Glu Ala Tyr Glu Arg 435 440 Leu Ile Leu Asp Ala Leu Leu Asp Glu Ser Ser Leu Phe Pro Thr Asn 455 Glu Glu Val Glu Leu Ser Trp Lys Ile Leu Asp Pro Ile Leu Glu Ala 470 Trp Asp Ala Asp Gly Glu Pro Glu Asp Tyr Pro Ala Gly Thr Trp Gly Pro Lys Ser Ala Asp Glu Met Leu Ser Arg Asn Gly His Thr Trp Arg 505 510 Arg Pro <210> 245 <211> 1203 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(1180) <223> RXA02738 <400> 245 ttgttgttaa tcggtacaaa gggtcttaag cacatccctt acttgcctgc tctccttgag 60 cacagttcaa gaacaattct tttaaggaaa atttagtttc atg tct cac att gat Met Ser His Ile Asp gat ctt gca cag ctc ggc act tcc act tgg ctc gac gac ctc tcc cgc 163 Asp Leu Ala Gln Leu Gly Thr Ser Thr Trp Leu Asp Asp Leu Ser Arg 211 gag cgc att act tcc ggc aat ctc agc cag gtt att gag gaa aag tct

Glu Arg Ile Thr Ser Gly Asn Leu Ser Gln Val Ile Glu Glu Lys Ser

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gac Asp	ggc Gly	cgc Arg	gtg Val 105	tcc Ser	atc Ile	gag Glu	gtt Val	gac Asp 110	cca Pro	cgt Arg	atc Ile	tct Ser	gct Ala 115	gac Asp	cgc Arg	451
							aag Lys 125									499
							cct Pro									547
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	-	-		-	-	-	gct Ala		_	-		_				835
							ggt Gly									883
_					-		cct Pro			-	-				_	931
tcc	gag	ctg	gct	ggt	cca	aac	acc	gtc	aac	acc	atg	cca	gaa	ggc	acc	979

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aac Asn 310	tcc Ser	gcg Ala	gca Ala	gaa Glu	gct Ala 315	gac Asp	gct Ala	gtg Val	ttc Phe	tcc Ser 320	cag Gln	ctt Leu	gag Glu	gct Ala	ctg Leu 325	1075
ggc Gly	gtt Val	gac Asp	ttg Leu	gca Ala 330	gat Asp	gtc Val	ttc Phe	cag Gln	gtc Val 335	ctg Leu	gag Glu	acc Thr	gag Glu	ggt Gly 340	gtg Val	1123
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Gly 145	Ser	Leu	Pro	Ala	Ile 150	Thr	Asp	Ala	Leu	Ala 155	Glu	Gly	Ile	Ser	Val 160	
Asn	Val	Thr	Leu	Ile	Phe	Ser	Val	Ala	Arg	Tyr	Arg	Glu	Val	Ile	Ala	

170 175 165 Ala Phe Ile Glu Gly Ile Lys Gln Ala Ala Ala Asn Gly His Asp Val 185 Ser Lys Ile His Ser Val Ala Ser Phe Phe Val Ser Arg Val Asp Val Glu Ile Asp Lys Arg Leu Glu Ala Ile Gly Ser Asp Glu Ala Leu Ala Leu Arg Gly Lys Ala Gly Val Ala Asn Ala Gln Arg Ala Tyr Ala Val 230 Tyr Lys Glu Leu Phe Asp Ala Ala Glu Leu Pro Glu Gly Ala Asn Thr Gln Arg Pro Leu Trp Ala Ser Thr Gly Val Lys Asn Pro Ala Tyr Ala Ala Thr Leu Tyr Val Ser Glu Leu Ala Gly Pro Asn Thr Val Asn Thr Met Pro Glu Gly Thr Ile Asp Ala Val Leu Glu Gln Gly Asn Leu His 295 Gly Asp Thr Leu Ser Asn Ser Ala Ala Glu Ala Asp Ala Val Phe Ser 310 315 Gln Leu Glu Ala Leu Gly Val Asp Leu Ala Asp Val Phe Gln Val Leu 325 Glu Thr Glu Gly Val Asp Lys Phe Val Ala Ser Trp Ser Glu Leu Leu 345 Glu Ser Met Glu Ala Arg Leu Lys 355 <210> 247 <211> 2223 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(2200) <223> RXA02739 <400> 247 cctttgccaa atttgaacca attaacctaa gtcgtagatc tgatcatcgg atctaacgaa 60 aacgaaccaa aactttggtc ccggtttaac ccaggaagga ttg acc acc ttg acg 115 Leu Thr Thr Leu Thr 1 163 ctg tca cct gaa ctt cag gcg ctc act gta cgc aat tac ccc tct gat Leu Ser Pro Glu Leu Gln Ala Leu Thr Val Arg Asn Tyr Pro Ser Asp 20 10

tgg tcc gat gtg gac acc aag gct gta gac act gtt cgt gtc ctc gct

211

Trp	Ser	Asp	Val 25	Asp	Thr	Lys	Ala	Val 30	Asp	Thr	Val	Arg	Val 35	Leu	Ala	
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-	-	-			gca Ala			_		_		_	_		_	307
					aac Asn 75											355
_					ttg Leu		-			-			-			403
					gat Asp											451
					cct Pro											499
					ggc Gly	-			-		_	_		-	-	547
					gag Glu 155											595
					gac Asp											643
					gtc Val											691
					ctc Leu											739
					gag Glu											787
	_	_			tgg Trp 235	-				-		-			_	835
					gct						aag Lys					883
	Ala	AIG		250	AIG	1124			255		-	-	•	260	-3-	

265 270 275

			aac Asn													979
			gca Ala													1027
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			cag Gln													1123
			aac Asn 345													1171
			cca Pro													1219
			ggc Gly													1267
-	_		aag Lys													1315
			aac Asn													1363
			tcc Ser 425													1411
			ggt Gly													1459
			cac His													1507
			tac Tyr													1555
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-			acc Thr 505		_		-	_		_	-	-	_	-	-	1651

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gca ctg acc cgc cag aac gtt cct gtt ctg gaa ggc acc aag gag aag Ala Leu Thr Arg Gln Asn Val Pro Val Leu Glu Gly Thr Lys Glu Lys 550 565 560 565	1795
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Val Arg Val Leu Ala Ala Asp Ala Val Glu Asn Cys Gly Ser Gly His Pro Gly Thr Ala Met Ser Leu Ala Pro Leu Ala Tyr Thr Leu Tyr Gln Arg Val Met Asn Val Asp Pro Gln Asp Thr Asn Trp Ala Gly Arg Asp Arg Phe Val Leu Ser Cys Gly His Ser Ser Leu Thr Gln Tyr Ile Gln Leu Tyr Leu Gly Gly Phe Gly Leu Glu Met Asp Asp Leu Lys Ala Leu Arg Thr Trp Asp Ser Leu Thr Pro Gly His Pro Glu Tyr Arg His Thr 120 Lys Gly Val Glu Ile Thr Thr Gly Pro Leu Gly Gln Gly Leu Ala Ser 135 Ala Val Gly Met Ala Met Ala Ala Arg Arg Glu Arg Gly Leu Phe Asp 150 155 Pro Thr Ala Ala Glu Gly Glu Ser Pro Phe Asp His His Ile Tyr Val 170 Ile Ala Ser Asp Gly Asp Leu Gln Glu Gly Val Thr Ser Glu Ala Ser 185 Ser Ile Ala Gly Thr Gln Gln Leu Gly Asn Leu Ile Val Phe Trp Asp Asp Asn Arg Ile Ser Ile Glu Asp Asn Thr Glu Ile Ala Phe Asn Glu Asp Val Val Ala Arg Tyr Lys Ala Tyr Gly Trp Gln Thr Ile Glu Val Glu Ala Gly Glu Asp Val Ala Ala Ile Glu Ala Ala Val Ala Glu Ala 250 Lys Lys Asp Thr Lys Arg Pro Thr Phe Ile Arg Val Arg Thr Ile Ile Gly Phe Pro Ala Pro Thr Met Met Asn Thr Gly Ala Val His Gly Ala 280 Ala Leu Gly Ala Ala Glu Val Ala Ala Thr Lys Thr Glu Leu Gly Phe Asp Pro Glu Ala His Phe Ala Ile Asp Asp Glu Val Ile Ala His Thr 315 310 Arg Ser Leu Ala Glu Arg Ala Ala Gln Lys Lys Ala Ala Trp Gln Val Lys Phe Asp Glu Trp Ala Ala Ala Asn Pro Glu Asn Lys Ala Leu Phe 340 345

Asp Arg Leu Asn Ser Arg Glu Leu Pro Ala Gly Tyr Ala Asp Glu Leu 360 Pro Thr Trp Asp Ala Asp Glu Lys Gly Val Ala Thr Arg Lys Ala Ser Glu Ala Ala Leu Gln Ala Leu Gly Lys Thr Leu Pro Glu Leu Trp Gly Gly Ser Ala Asp Leu Ala Gly Ser Asn Asn Thr Val Ile Lys Gly Ser Pro Ser Phe Gly Pro Glu Ser Ile Ser Thr Glu Thr Trp Ser Ala Glu Pro Tyr Gly Arg Asn Leu His Phe Gly Ile Arg Glu His Ala Met Gly Ser Ile Leu Asn Gly Ile Ser Leu His Gly Gly Thr Arg Pro Tyr Gly Gly Thr Phe Leu Ile Phe Ser Asp Tyr Met Arg Pro Ala Val Arg Leu 475 Ala Ala Leu Met Glu Thr Asp Ala Tyr Tyr Val Trp Thr His Asp Ser Ile Gly Leu Gly Glu Asp Gly Pro Thr His Gln Pro Val Glu Thr Leu 500 505 Ala Ala Leu Arg Ala Ile Pro Gly Leu Ser Val Leu Arg Pro Ala Asp 520 Ala Asn Glu Thr Ala Gln Ala Trp Ala Ala Ala Leu Glu Tyr Lys Glu Gly Pro Lys Gly Leu Ala Leu Thr Arg Gln Asn Val Pro Val Leu Glu Gly Thr Lys Glu Lys Ala Ala Glu Gly Val Arg Arg Gly Gly Tyr Val Leu Val Glu Gly Ser Lys Glu Thr Pro Asp Val Ile Leu Met Gly Ser Gly Ser Glu Val Gln Leu Ala Val Asn Ala Ala Lys Ala Leu Glu Ala Glu Gly Val Ala Ala Arg Val Val Ser Val Pro Cys Met Asp Trp Phe 615 Gln Glu Gln Asp Ala Glu Tyr Ile Glu Ser Val Leu Pro Ala Ala Val Thr Ala Arg Val Ser Val Glu Ala Gly Ile Ala Met Pro Trp Tyr Arg 650 Phe Leu Gly Thr Gln Gly Arg Ala Val Ser Leu Glu His Phe Gly Ala Ser Ala Asp Tyr Gln Thr Leu Phe Glu Lys Phe Gly Ile Thr Thr Asp

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Phe His Asp Leu Pro Leu Glu Glu Arg Leu Thr Leu Ala Arg Leu Gly
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Thr Ser His Tyr Ser Arg Gln Leu Ser Leu Val Asp Asn Ala Glu Phe
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gag gaa att gcc tac ggt tcc acg ctc aat ccc gat gcg ttg cgt aac 344 Glu Glu Ile Ala Tyr Gly Ser Thr Leu Asn Pro Asp Ala Leu Arg Asn 85 90 95

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Val Pro Ala Ser Glu Thr Leu Trp Met Arg Ser Arg Glu Val Trp Ile
135 140 145

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Val 145	Trp	Ile	His	Ala	Val 150	Asp	Leu	Gly	Ala	Val 155	Ala	Thr	Phe	Gly	Asp 160	
Ile	Pro	Glu	Val	Ile 165	Leu	Arg	Thr	Leu	Ala 170	Ala	Glu	Ile	Thr	Gln 175	Lys	

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	ggc Gly 135	-	-		-						_			 547
	gca Ala													595
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-	cag Gln	-				-				_		_	_	 739
	cag Gln 215													787
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	gat Asp													883
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	tcc Ser 295													1027
	ggt Gly													1075
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ctc ggc gac Leu Gly Asp		Asp Se										1363
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gac agc ctg Asp Ser Leu 440	Arg Ala											1459
cgc gac ttc Arg Asp Phe 455			Thr									1507
tcc ttc cac Ser Phe His		Trp Se										1552
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<pre>taaaggctct  &lt;210&gt; 252 &lt;211&gt; 484 &lt;212&gt; PRT &lt;213&gt; Coryn  &lt;400&gt; 252 Met Thr Asn 1</pre>	ebacteri Gly Asp 5 Ser Asn 20 Tyr Asn	ca caa um gluta Asn Leu Leu Ala	a Ala a Arg	Gln Asn 25	10 Phe	Gly Ala	Arg	Asn	Gly 30	15 Asn	Thr	1575
<pre>taaaggctct  &lt;210&gt; 252 &lt;211&gt; 484 &lt;212&gt; PRT &lt;213&gt; Coryn &lt;400&gt; 252 Met Thr Asn 1  Val Met Gly  Val Ala Val</pre>	ebacteri Gly Asp 5 Ser Asn 20 Tyr Asn	um gluta Asn Leu Leu Ala Arg Se	Ala Arg Thr 40	Gln Asn 25 Asp	10 Phe Lys	Gly Ala Thr	Arg Asp	Asn Lys 45	Gly 30 Leu	15 Asn Ile	Thr Ala	1575
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Thr	Ile	Arg 115	Arg	Glu	Lys	Glu	Ile 120	Ser	Ala	Arg	Gly	Leu 125	His	Phe	Val
Gly	Ala 130	Gly	Ile	Ser	Gly	Gly 135	Glu	Glu	Gly	Ala	Leu 140	Asn	Gly	Pro	Ser
Ile 145	Met	Pro	Gly	Gly	Pro 150	Ala	Lys	Ser	Tyr	Glu 155	Ser	Leu	Gly	Pro	Leu 160
Leu	Glu	Ser	Ile	Ala 165	Ala	Asn	Val	Asp	Gly 170	Thr	Pro	Суѕ	Val	Thr 175	His
Ile	Gly	Pro	Asp 180	Gly	Ala	Gly	His	Phe 185	Val	Lys	Met	Val	His 190	Asn	Gly
Ile	Glu	Tyr 195	Ala	Asp	Met	Gln	Val 200	Ile	Gly	Glu	Ala	Tyr 205	His	Leu	Leu
Arg	Tyr 210	Ala	Ala	Gly	Met	Gln 215	Pro	Ala	Glu	Ile	Ala 220	Glu	Val	Phe	Lys
Glu 225	Trp	Asn	Ala	Gly	Asp 230	Leu	Asp	Ser	Туr	Leu 235	Ile	Glu	Ile	Thr	Ala 240
Glu	Val	Leu	Ser	Gln 245	Val	Asp	Ala	Glu	Thr 250	Gly	Lys	Pro	Leu	Ile 255	Asp
Val	Ile	Val	Asp 260	Ala	Ala	Gly	Gln	Lys 265	Gly	Thr	Gly	Arg	Trp 270	Thr	Val
Lys	Ala	Ala 275	Leu	Asp	Leu	Gly	Ile 280	Ala	Thr	Thr	Gly	Ile 285	Gly	Glu	Ala
Val	Phe 290	Ala	Arg	Ala	Leu	Ser 295	Gly	Ala	Thr	Ser	Gln 300	Arg	Ala	Ala	Ala
Gln 305	Gly	Asn	Leu	Pro	Ala 310	Gly	Val	Leu	Thr	Asp 315	Leu	Glu	Ala	Leu	Gly 320
Val	Asp	Lys	Ala	Gln 325	Phe	Val	Glu	Asp	Val 330	Arg	Arg	Ala	Leu	Tyr 335	Ala
Ser	Lys	Leu	Val 340	Ala	Tyr	Ala	Gln	Gly 345	Phe	Asp	Glu	Ile	Lys 350	Ala	Gly
Ser	Asp	Glu 355	Asn	Asn	Trp	Asp	Val 360	Asp	Pro	Arg	Asp	Leu 365	Ala	Thr	Ile
Trp	Arg 370	Gly	Gly	Суз	Ile	Ile 375	Arg	Ala	Lys	Phe	Leu 380	Asn	Arg	Ile	Val
Gľu 385	Ala	Tyr	Asp	Ala	Asn 390	Ala	Glu	Leu	Glu	Ser 395	Leu	Leu	Leu	Asp	Pro 400
Tyr	Phe	Lys	Ser	Glu 405	Leu	Gly	Asp	Leu	Ile 410	Asp	Ser	Trp	Arg	Arg 415	Val
Ile	Val	Thr	Ala 420	Thr	Gln	Leu	Gly	Leu 425	Pro	Ile	Pro	Val	Phe 430	Ala	Ser

Ser Leu Ser Tyr Tyr Asp Ser Leu Arg Ala Glu Arg Leu Pro Ala Ala Leu Ile Gln Gly Gln Arg Asp Phe Phe Gly Ala His Thr Tyr Lys Arg Ile Asp Lys Asp Gly Ser Phe His Thr Glu Trp Ser Gly Asp Arg Ser Glu Val Glu Ala <210> 253 <211> 1537 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(1537) <223> FRXA00999 <400> 253 cctcctgtga cctggtaaaa tcgccactac ccccaaatgg tcacaccttt taggccgatt 60 ttgctgacac cgggctatgc cgtcaagtac gatcaataac atg act aat gga gat 115 Met Thr Asn Gly Asp 1 aat ctc qca cag atc ggc gtt gta ggc cta gca gta atg ggc tca aac Asn Leu Ala Gln Ile Gly Val Val Gly Leu Ala Val Met Gly Ser Asn 10 ctc gcc cgc aac ttc gcc cgc aac ggc aac act gtc gct gtc tac aac 211 Leu Ala Arg Asn Phe Ala Arg Asn Gly Asn Thr Val Ala Val Tyr Asn 25 cgc agc act gac aaa acc gac aag ctc atc gcc gat cac ggc tcc gaa 259 Arg Ser Thr Asp Lys Thr Asp Lys Leu Ile Ala Asp His Gly Ser Glu 40 45 307 ggc aac ttc atc cct tct gca acc gtc gaa gag ttc gta gca tcc ctg Gly Asn Phe Ile Pro Ser Ala Thr Val Glu Glu Phe Val Ala Ser Leu 55 gaa aag cca cgc cgc gcc atc atc atg gtt cag gct ggt aac gcc acc 355 Glu Lys Pro Arg Arg Ala Ile Ile Met Val Gln Ala Gly Asn Ala Thr 70 gac gca gtc atc aac cag ctg gca gat gcc atg gac gaa ggc gac atc 403 Asp Ala Val Ile Asn Gln Leu Ala Asp Ala Met Asp Glu Gly Asp Ile 90 ate ate gae gge gge aac gee ete tae ace gae ace att egt ege gag 451 Ile Ile Asp Gly Gly Asn Ala Leu Tyr Thr Asp Thr Ile Arg Arg Glu 105 aag gaa atc tcc gca cgc ggt ctc cac ttc gtc ggt gct ggt atc tcc Lys Glu Ile Ser Ala Arg Gly Leu His Phe Val Gly Ala Gly Ile Ser 120

ggc ggc g Gly Gly G 135	gaa gaa go Glu Glu G	gc gca cto ly Ala Leo 140	Asn Gly	cca tcc Pro Ser	atc atg Ile Met 145	cct ggt Pro Gly	ggc 54 Gly	<b>1</b> 7
cca gca a Pro Ala L 150	ag too ta Lys Ser T	ac gag tco yr Glu Se: 155	ctc gga Leu Gly	cca ctg Pro Leu 160	ctt gag Leu Glu	tcc atc Ser Ile	gct 59 Ala 165	95
gcc aac g Ala Asn V	/al Asp G	gc acc cca ly Thr Pro 70	tgt gto Cys Val	acc cac Thr His 175	atc ggc Ile Gly	cca gac Pro Asp 180	ggc 64 Gly	43
gcc ggc c Ala Gly H	cac ttc g His Phe Va 185	tc aag ato al Lys Me	gtc cad Val His	Asn Gly	atc gag Ile Glu	tac gcc Tyr Ala 195	gac 69 Asp	91
Met Gln V		gc gag gca ly Glu Ala						39
		aa atc gc lu Ile Ala 22	a Glu Val				-	87
		ac ctc ato yr Leu Ilo 235					•	35
	Ala Glu T	cc ggc aachr Gly Ly: 50						83
		gc acc gg ly Thr Gl		Thr Val				31
Leu Gly I		cc acc gg hr Thr Gl						79
	Sly Ala T	cc agc cachr Ser Gla	Arg Ala	Ala Ala	Gln Gly			027
		cc gat ct hr Asp Le 315						075
	Glu Asp V	tt cgc cg al Arg Ar 30					-	123
		tc gac ga he Asp Gl		Ala Gly				171
Trp Asp V		ct cgc ga ro Arg As	Leu Ala				- 5	219

Ile	att Ile 375	cgc Arg	gct Ala	aag Lys	ttc Phe	ctc Leu 380	aac Asn	cgc Arg	atc Ile	gtc Val	gaa Glu 385	gca Ala	tac Tyr	gat Asp	gca Ala	1267
					tcc Ser 395											1315
					gat Asp											1363
					atc Ile											1411
gac Asp	agc Ser	ctg Leu 440	cgt Arg	gca Ala	gag Glu	cgt Arg	ctg Leu 445	cca Pro	gca Ala	gcc Ala	ctg Leu	atc Ile 450	caa Gln	gga Gly	cag Gln	1459
					gcg Ala											1507
					tgg Trp 475			-	_							1537
	0> 2! 1> 4					· i					٠					
	2> PI 3> Co		ebact	eri	ım gi	lutar	nicur	n								
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<21: <400 Met 1 Val Val Asp Phe 65 Ala Asp	3> Co 0> 2! Thr Met Ala His 50 Val Gly	Gly Val 35 Gly Ala Asn	Ser 20 Tyr Ser Ala Asp	Asp 5 Asn Asn Glu Leu Thr 85 Ile	Asn Leu Arg Gly Glu 70 Asp	Leu Ala Ser Asn 55 Lys Ala	Ala Arg Thr 40 Phe Pro Val	Gln Asn 25 Asp Ile Arg Ile Gly 105	10 Phe Lys Pro Arg Asn 90 Gly	Ala Thr Ser Ala 75 Gln Asn	Arg Asp Ala 60 Ile Leu Ala	Asn Lys 45 Thr Ile Ala	Gly 30 Leu Val Met Asp	15 Asn Ile Glu Val Ala 95 Thr	Thr Ala Glu Gln 80 Met	

Ile Met Pro Gly Gly Pro Ala Lys Ser Tyr Glu Ser Leu Gly Pro Leu Leu Glu Ser Ile Ala Ala Asn Val Asp Gly Thr Pro Cys Val Thr His Ile Gly Pro Asp Gly Ala Gly His Phe Val Lys Met Val His Asn Gly Ile Glu Tyr Ala Asp Met Gln Val Ile Gly Glu Ala Tyr His Leu Leu Arg Tyr Ala Ala Gly Met Gln Pro Ala Glu Ile Ala Glu Val Phe Lys Glu Trp Asn Ala Gly Asp Leu Asp Ser Tyr Leu Ile Glu Ile Thr Ala Glu Val Leu Ser Gln Val Asp Ala Glu Thr Gly Lys Pro Leu Ile Asp Val Ile Val Asp Ala Ala Gly Gln Lys Gly Thr Gly Arg Trp Thr Val Lys Ala Ala Leu Asp Leu Gly Ile Ala Thr Thr Gly Ile Gly Glu Ala 280 Val Phe Ala Arg Ala Leu Ser Gly Ala Thr Ser Gln Arg Ala Ala Ala Gln Gly Asn Leu Pro Ala Gly Val Leu Thr Asp Leu Glu Ala Leu Gly 315 310 Val Asp Lys Ala Gln Phe Val Glu Asp Val Arg Arg Ala Leu Tyr Ala Ser Lys Leu Val Ala Tyr Ala Gln Gly Phe Asp Glu Ile Lys Ala Gly Ser Asp Glu Asn Asn Trp Asp Val Asp Pro Arg Asp Leu Ala Thr Ile 360 Trp Arg Gly Gly Cys Ile Ile Arg Ala Lys Phe Leu Asn Arg Ile Val Glu Ala Tyr Asp Ala Asn Ala Glu Leu Glu Ser Leu Leu Leu Asp Pro 390 395 Tyr Phe Lys Ser Glu Leu Gly Asp Leu Ile Asp Ser Trp Arg Arg Val Ile Val Thr Ala Thr Gln Leu Gly Leu Pro Ile Pro Val Phe Ala Ser 425 Ser Leu Ser Tyr Tyr Asp Ser Leu Arg Ala Glu Arg Leu Pro Ala Ala 435 Leu Ile Gln Gly Gln Arg Asp Phe Phe Gly Ala His Thr Tyr Lys Arg 455 450 460

Ile Asp Lys Asp Gly Ser Phe His Thr Glu Trp Ser Gly Asp Arg

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160

155

cag act gat cc Gln Thr Asp Pr					
gtt cgc tac aa Val Arg Tyr As 18	n Phe Asn	•	•	_	
ctt ccc aca ga Leu Pro Thr As 200	c ggc tac o p Gly Tyr i	gcg gca tgg Ala Ala Trp 205	ttg gaa aag Leu Glu Lys	atg gca gag Met Ala Glu 210	cat 739 His
gag ctt atc ga Glu Leu Ile As 215	p Val Arg	-			-
gac ctc cgc gc Asp Leu Arg Al 230	•				
ctc gac ctc ta Leu Asp Leu Ty					
ctc gac ttt ga Leu Asp Phe Gl 26	u Thr Glu				
cca gtg atg aa Pro Val Met As 280			-	-	
gag ttc cgt ca Glu Phe Arg Hi 295	s Phe His				
aag acc gtc at Lys Thr Val Il 310					
gag cct tat ta Glu Pro Tyr Ty		Asn Thr Pro		Asp Met Leu	Lys
cag tac cgc ct Gln Tyr Arg Le 34	u Leu Ala i				
ttc ggc ggt cg Phe Gly Gly Ar 360					
atc ggt tct gc Ile Gly Ser Al 375	a Leu Ser 1				
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ggcatctccc aca					1326

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<211> 401

<212> PRT

<213> Corynebacterium glutamicum

<400> 256

Met Thr Glu Ser Lys Asn Tyr Asp Leu Ile Val Val Gly Ser Gly Leu
1 5 10 15

Phe Gly Leu Thr Val Ala Glu Arg Ala Ala Ser Gln Leu Gly Lys Lys 20 25 30

Val Leu Ile Val Glu Arg Arg Ser His Leu Gly Gly Asn Ala Tyr Ser 35 40 45

Glu Ala Glu Pro Glu Thr Gly Ile Glu Ile His Lys Tyr Gly Ala His 50 55 60

Leu Phe His Thr Ser Asn Thr Arg Val Trp Glu Tyr Val Asn Gln Phe 65 70 75 80

Thr Ser Phe Thr Gly Tyr Gln His Arg Val Phe Ala Met His Asn Gly 85 90 95

Thr Ala Tyr Gln Phe Pro Met Gly Leu Gly Leu Ile Asn Gln Phe Phe 100 105 110

Gly Lys Tyr Tyr Ser Pro Asp Glu Ala Arg Glu Leu Ile Lys Glu Gln 115 120 125

Ser Ala Glu Ile Asp Ser Ser Asp Ala Thr Asn Leu Glu Glu Lys Ala 130 135 140

Ile Ser Leu Ile Gly Arg Pro Leu Tyr Glu Ala Phe Ile Arg Asp Tyr 145 150 155 160

Thr Ala Lys Gln Trp Gln Thr Asp Pro Lys Asn Leu Pro Ala Gly Asn 165 170 175

Ile Thr Arg Leu Pro Val Arg Tyr Asn Phe Asn Asn Arg Tyr Phe Asn 180 185 190

Asp Thr Tyr Glu Gly Leu Pro Thr Asp Gly Tyr Ala Ala Trp Leu Glu 195 200 205

Lys Met Ala Glu His Glu Leu Ile Asp Val Arg Leu Asp Thr Asp Trp 210 215 220

Phe Asp Val Arg Asp Asp Leu Arg Ala Ser Asn Pro Asp Ala Pro Val 225 230 235 240

Val Tyr Thr Gly Pro Leu Asp Leu Tyr Phe Asn Tyr Ala Glu Gly Lys 245 250 255

Leu Gly Trp Arg Thr Leu Asp Phe Glu Thr Glu Val Val Glu Thr Gly 260 265 270

Asp Phe Gln Gly Thr Pro Val Met Asn Tyr Asn Asp Ala Asp Val Pro 275 280 285

Phe Thr Arg Ile His Glu Phe Arg His Phe His Pro Glu Arg Asp Asp 290 295 Ser Tyr Pro Lys Asp Lys Thr Val Ile Met Arg Glu Phe Ser Arg Phe 315 310 Ala Asp Asn Glu Asp Glu Pro Tyr Tyr Pro Ile Asn Thr Pro Asp Asp 325 330 Arg Asp Met Leu Lys Gln Tyr Arg Leu Leu Ala Ala Glu Glu Ala Ala 345 Asn Asn Lys Val Leu Phe Gly Gly Arg Leu Gly Thr Tyr Gln Tyr Leu 360 365 355 Asp Met His Met Ala Ile Gly Ser Ala Leu Ser Met Phe Asp Asn Lys 375 Leu Val Pro Phe Phe Glu Glu Gly Thr Pro Leu Glu Gln Glu Arg Gly 400 His <210> 257 <211> 512 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (1)..(489) <223> FRXA02596 <400> 257 cct gtg gtc tac acc ggc cca ctc gac ctc tac ttc aac tac gca gag 48 Pro Val Val Tyr Thr Gly Pro Leu Asp Leu Tyr Phe Asn Tyr Ala Glu ggc aag ctg gga tgg cgc acc ctc gac ttt gaa acc gaa gta gta gaa 96 Gly Lys Leu Gly Trp Arg Thr Leu Asp Phe Glu Thr Glu Val Val Glu acc ggt gac ttc caa gga acc cca gtg atg aac tac aac gat gcg gac 144 Thr Gly Asp Phe Gln Gly Thr Pro Val Met Asn Tyr Asn Asp Ala Asp 40 gta ect tte ace ege ate cae gag tte egt cae tte cae eca gag egt 192 Val Pro Phe Thr Arg Ile His Glu Phe Arg His Phe His Pro Glu Arg 55 gat gac agt tac ccc aag gat aag acc gtc atc atg cgc gag ttc tcc 240 Asp Asp Ser Tyr Pro Lys Asp Lys Thr Val Ile Met Arg Glu Phe Ser 70 cgt ttc gca gat aac gag gat gag cct tat tac cca atc aac act cca 288 Arg Phe Ala Asp Asn Glu Asp Glu Pro Tyr Tyr Pro Ile Asn Thr Pro 85 90

							cag Gln						gag Glu	336
							ttc Phe 120							384
							atc Ile						gac Asp	432
	-	-		•			gaa Glu	-		-	_	_	_	480
_	gga Gly		taaa	aagga	aag q	ggcat	tete	cc ac	ca					512

<210> 258

<211> 163

<212> PRT

<213> Corynebacterium glutamicum

<400> 258

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Gly Lys Leu Gly Trp Arg Thr Leu Asp Phe Glu Thr Glu Val Val Glu 20 25 30

Thr Gly Asp Phe Gln Gly Thr Pro Val Met Asn Tyr Asn Asp Ala Asp 35 40 45

Val Pro Phe Thr Arg Ile His Glu Phe Arg His Phe His Pro Glu Arg 50 55 60

Asp Asp Ser Tyr Pro Lys Asp Lys Thr Val Ile Met Arg Glu Phe Ser 65 70 75 80

Arg Phe Ala Asp Asn Glu Asp Glu Pro Tyr Tyr Pro Ile Asn Thr Pro 85 90 95

Asp Asp Arg Asp Met Leu Lys Gln Tyr Arg Leu Leu Ala Ala Glu Glu 100 105 110

Ala Ala Asn Asn Lys Val Leu Phe Gly Gly Arg Leu Gly Thr Tyr Gln 115 120 125

Tyr Leu Asp Met His Met Ala Ile Gly Ser Ala Leu Ser Met Phe Asp 130 135 140

Asn Lys Leu Val Pro Phe Phe Glu Glu Gly Thr Pro Leu Glu Gln Glu 145 150 155 160

Arg Gly His

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<210> 260 <211> 166 <212> PRT <213> Corynebacterium glutamicum Met Thr Glu Ser Lys Asn Tyr Asp Leu Ile Val Val Gly Ser Gly Leu Phe Gly Leu Thr Val Ala Glu Arg Ala Ala Ser Gln Leu Gly Lys Lys Val Leu Ile Val Glu Arg Arg Ser His Leu Gly Gly Asn Ala Tyr Ser 40 Glu Ala Glu Pro Glu Thr Gly Ile Glu Ile His Lys Tyr Gly Ala His Leu Phe His Thr Ser Asn Thr Arg Val Trp Glu Tyr Val Asn Gln Phe 75 Thr Ser Phe Thr Gly Tyr Gln His Arg Val Phe Ala Met His Asn Gly Thr Ala Tyr Gln Phe Pro Met Gly Leu Gly Leu Ile Asn Gln Phe Phe Gly Lys Tyr Tyr Ser Pro Asp Glu Ala Arg Glu Leu Ile Lys Glu Gln Ser Ala Glu Ile Asp Ser Ser Asp Ala Thr Asn Leu Glu Glu Lys Ala Ile Ser Leu Ile Gly Arg Pro Leu Tyr Glu Ala Phe Ile Arg Asp Tyr 155 Thr Ala Lys Gln Trp Gln <210> 261 <211> 668 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (1)..(645) <223> RXA02572 <400> 261 gcg gtc gct gag att tgc gag ccg acc ggc gcc gat gcg gtt gcg ctt Ala Val Ala Glu Ile Cys Glu Pro Thr Gly Ala Asp Ala Val Ala Leu gtg gat gcc atc ggt cac gac gat cgt atc ggc cga aag ttc tta ggc Val Asp Ala Ile Gly His Asp Asp Arg Ile Gly Arg Lys Phe Leu Gly 20 gcg ggc ctg gga ttc ggt ggc ggt tgt ttg cct aaa gac atc cgc gct

Ala Gly Leu 35		Gly Gly	Gly Cys 40	Leu Pro		p Ile A 5	arg Ala	
ttc atg gca Phe Met Ala 50	cgc gcg Arg Ala	ggc gaa Gly Glu 55	ttg ggt Leu Gly	gct gad Ala Asp	c cag go o Gln Al 60	a tta a a Leu T	cg ttc hr Phe	192
ttg cgt gag Leu Arg Glu 65					g Arg As			240
cag ctg gcc Gln Leu Ala								288
aca gtg ctc Thr Val Leu								336
tct ccg gcg Ser Pro Ala 115	Leu Ser					n Gly A		384
gtc tcg gtc Val Ser Val 130	tac gac Tyr Asp	ccg gaa Pro Glu 135	gct atg Ala Met	gac aad Asp Asi	c gct cg n Ala Ar 140	a cgc g g Arg V	tc ttc al Phe	432
ccg acg ctc Pro Thr Leu 145					u Ala Le			480
cac ctc gtc His Leu Val						g Asp L		528
ccc gaa gtg Pro Glu Val				Lys Arc				576
cga aac gtc Arg Asn Val 195	Leu Asp					y Trp G		624
gaa gcg ctc Glu Ala Leu 210			tagtgcg	gtg gato	caggcgg	ggc		668
<210> 262 <211> 215 <212> PRT <213> Coryn	ebacteriu	ım glutan	nicum					
<400> 262								
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Val Asp Ala	Ile Gly 20	His Asp	Asp Arg 25		y Arg Ly	s Phe L 30	eu Gly	
Ala Gly Leu	Gly Phe	Gly Gly	Gly Cys	Leu Pro	Lys As	p Ile A	rg Ala	

		35					40					45				
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Leu 65	Arg	Glu	Val	Asp	Ser 70	Ile	Asn	Met	Arg	Arg 75	Arg	Asp	Arg	Val	Val 80	
Gln	Leu	Ala	Lys	Glu 85	Met	Cys	Gly	Gly	Ser 90	Leu	Leu	Gly	Lys	Arg 95	Val	
Thr	Val	Leu	Gly 100	Ala	Ala	Phe	Lys	Pro 105	Asn	Ser	Asp	Asp	Val 110	Arg	Asp	
Ser	Pro	Ala 115	Leu	Ser	Val	Ala	Gly 120	Ser	Leu	Ser	Leu	Gln 125	Gly	Ala	Ala	
Val	Ser 130	Val	Tyr	Asp	Pro	Glu 135	Ala	Met	Asp	Asn	Ala 140	Arg	Arg	Val	Phe	
Pro 145	Thr	Leu	Ser	Tyr	Ala 150	Ser	Ser	Thr	Lys	Glu 155	Ala	Leu	Ile	Asp	Ala 160	
His	Leu	Val	Val	Leu 165	Ala	Thr	Glu	Trp	Gln 170	Glu	Phe	Arg	Asp	Leu 175	Asp	
Pro	Glu	Val	Ala 180	Gly	Gly	Val	Val	Glu 185	Lys	Arg	Ala	Ile	Ile 190	Asp	Gly	
Arg	Asn	Val 195	Leu	Asp	Val	Ala	Lys 200	Trp	Lys	Ala	Ala	Gly 205	Trp	Glu	Met ·	
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tac	gcgti	tgc (	catg	aggat	ta aq	gacta	accgt	t <b>t</b> aq	gtgg	ggtg		gat Asp				115
				gcc Ala												163

211

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	gtc Val															355
	gac Asp															403
	ggc Gly															451
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	cca Pro 135															547
	acc Thr															595
	gac Asp															643
	acc Thr															691
	gga Gly		Ser	Ala	Pro	Leu		Phe	Gly	Glu	Leu		Arg			739
	atc Ile 215															787
	gac Asp															835
	acc Thr															883
	gtc Val	-		-		_	-	-	-		-		-		-	931
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Leu Asp Ser Ser Leu Ala Gln Glu Ile Ala Ala Ile Asp Gly Val Glu  1 5 10 15  Leu Asp Ser Glu Val Thr Phe Ala Asp Leu Thr Thr Leu Arg Ile Gly	
Leu Asp Ser Ser Leu Ala Gln Glu Ile Ala Ala Ile Asp Gly Val Glu 1 10 15  Leu Asp Ser Glu Val Thr Phe Ala Asp Leu Thr Thr Leu Arg Ile Gly 20 25 30  Gly Lys Pro Arg Ser Ala Val Arg Cys Gln Thr Thr Glu Ala Leu Val	
Leu Asp Ser Ser Leu Ala Gln Glu Ile Ala Ala Ile Asp Gly Val Glu 15  Leu Asp Ser Glu Val Thr Phe Ala Asp Leu Thr Thr Leu Arg Ile Gly 25  Gly Lys Pro Arg Ser Ala Val Arg Cys Gln Thr Thr Glu Ala Leu Val 35  Ser Ala Ile Lys Leu Leu Asp Asp Ala Ser Leu Pro Leu Leu Ile Val	
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Leu Asp Ser Ser Leu Ala Gln Glu Ile Ala Ala Ile Asp Gly Val Glu 1 Ser Ala Ser Glu Val Thr Phe Ala Asp Leu Thr Thr Leu Arg Ile Gly 30 Ser Ala Ile Lys Leu Leu Asp Asp Ala Ser Leu Pro Leu Leu Ile Val 65 Gly Gly Gly Ser Asn Leu Val 70 Val Val Ala Asp Gly Asp Leu Asp Val Ile 65 Ala Val Ile Ile Glu Thr Asp Asp Val Ser Ile Asn Leu Thr Asp Gly 85 Ser Val Asp Ala Gly Leu Gly Gly Ile Glu Cys Leu Ser Gly Ile Pro	

His Gln Val Ser Trp Val Asp Ala Ser Glu Leu Asp Leu Ser Tyr Arg Tyr Ser Asn Leu Lys Phe Thr Asn Arg Ala Val Val Leu Ala Ile Glu 185 Leu Gln Leu Leu Thr Asp Gly Leu Ser Ala Pro Leu Arg Phe Gly Glu 200 Leu Gly Arg Arg Leu Ala Ile Ser Glu Ala Glu Pro His Pro Arg Arg Pro Val Arg Met Val Arg Asp Ala Val Leu Glu Leu Arg Arg Ala Lys 230 235 Gly Met Val Val Glu His Thr Asp His Asp Thr Trp Ser Ala Gly Ser Phe Phe Thr Asn Pro Ile Val Asp Pro Ala Leu Ala Asp Ala Val Phe 265 270 260 Glu Lys Val Gly Glu Pro Thr Met Pro Arg Phe Pro Ala Gly Asp Gly 280 Lys Glu Lys Leu Ser Ala Ala Trp Leu Ile Glu Arg Ala Gly Phe Lys Lys Gly His Pro Gly Ala Gly Ala Lys Ala Ser Leu Ser Thr Lys His Thr Leu Ala Leu Thr Asn Arg Gly Asp Ala Arg Ala Ser Asp Leu Val Ala Leu Ala Lys Glu Ile Arg Asp Gly Val Leu Glu Thr Phe Gly Val Thr Leu Val Pro Glu Pro Val Trp Ile Gly Ile Ser Ile Asp Asp 360 <210> 265 <211> 1124 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (1)..(1101) <223> RXA01216 <400> 265 acc gac cac act ctg tct gca ctg ctg gat gca cac gtg gaa gtt cca 48 Thr Asp His Thr Leu Ser Ala Leu Leu Asp Ala His Val Glu Val Pro acc gct gtc acc gtg ttg acc atg cgt ctg gat gac ccc acc ggc tac 96 Thr Ala Val Thr Val Leu Thr Met Arg Leu Asp Asp Pro Thr Gly Tyr 20 ggc cgc atc gtg cgc aac gaa gac gaa gtc acc gcc atc gtt gag

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	ggc Gly 130															432
	gca Ala															480
	atc Ile															528
	act Thr															576
	aac Asn	-				-		-		-		-				624
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	gaa Glu															768
	acc Thr															816
	tcc Ser		-					-		-		_				864

275 280 285 acc atc qgc agc cac gtt cgc act ggt tct gac acc atg ttt atc gct 912 Thr Ile Gly Ser His Val Arg Thr Gly Ser Asp Thr Met Phe Ile Ala 295 290 960 cca gtg acc gtg ggt gac gga gcg tat tcc gga gcc ggt aca gta att Pro Val Thr Val Gly Asp Gly Ala Tyr Ser Gly Ala Gly Thr Val Ile 310 1008 aaa gac gat gtt ccg cca gga gcc ctt gcc gtg tcc ggc gga cgc caa Lys Asp Asp Val Pro Pro Gly Ala Leu Ala Val Ser Gly Gly Arg Gln cga aac atc gaa ggc tgg gtg caa aag aag cgc cct gga acc gct gca 1056 Arg Asn Ile Glu Gly Trp Val Gln Lys Lys Arg Pro Gly Thr Ala Ala 1101 gca caa gcc gca gaa gcc gcc caa aac gtc cac aac cag gaa ggc Ala Gln Ala Ala Glu Ala Ala Gln Asn Val His Asn Gln Glu Gly taagcaggat cctcatgact gct 1124 <210> 266 <211> 367 <212> PRT <213> Corynebacterium glutamicum <400> 266 Thr Asp His Thr Leu Ser Ala Leu Leu Asp Ala His Val Glu Val Pro 5 10 Thr Ala Val Thr Val Leu Thr Met Arg Leu Asp Asp Pro Thr Gly Tyr Gly Arg Ile Val Arg Asn Glu Glu Gly Glu Val Thr Ala Ile Val Glu Gln Lys Asp Ala Ser Ala Glu Val Gln Ala Ile Asp Glu Val Asn Ser Gly Val Phe Ala Phe Asp Ala Ala Ile Leu Arg Ser Ala Leu Ala Glu Leu Lys Ser Asp Asn Ala Gln Gly Glu Leu Tyr Leu Thr Asp Val Leu Gly Ile Ala Arg Gly Glu Gly His Pro Val Arg Ala His Thr Ala Ala Asp Ala Arg Glu Leu Ala Gly Val Asn Asp Arg Val Gln Leu Ala Glu 120 Ala Gly Ala Glu Leu Asn Arg Arg Thr Val Ile Ala Ala Met Arg Gly Gly Ala Thr Ile Val Asp Pro Ala Thr Thr Trp Ile Asp Val Glu Val 145 150 155

ser	116	GIY	AIG	165	vai	116	116	птэ	170	GIY	1111	GIII	пец	175	GIY	
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Thr	Asn	Met 195	Thr	Ile	Gly	Asp	Gly 200	Ala	Ser	Val	Ile	Arg 205	Thr	His	Gly	
Phe	Asp 210	Ser	Thr	Ile	Gly	Glu 215	Asn	Ala	Thr	Val	Gly 220	Pro	Phe	Thr	Tyr	
Ile 225	Arg	Pro	Gly	Thr	Thr 230	Leu	Gly	Pro	Glu	Gly 235	Lys	Leu	Gly	Gly	Phe 240	
Val	Glu	Thr	Lys	Lys 245	Ala	Thr	Ile	Gly	Arg 250	Gly	Ser	Lys	Val	Pro 255	His	
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Ala	Ser	Ser 275	Val	Phe	Val	Asn	Tyr 280	Asp	Gly	Glu	Asn	Lys 285	His	His	Thr	
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Pro 305	Val	Thr	Val	Gly	Asp 310	Gly	Ala	Tyr	Ser	Gly 315	Ala	Gly	Thr	Val	Ile 320	
Lys	Asp	Asp	Val	Pro 325	Pro	Gly	Ala	Leu	Ala 330	Val	Ser	Gly	Gly	Arg 335	Gln	
Arg	Asn	Ile	Glu 340	Gly	Trp	Val	Gln	Lys 345	Lys	Arg	Pro	Gly	Thr 350	Ala	Ala	
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					aca Thr											163

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_		-		-			-	_	-	-	cga Arg	-	-		_	355
											ctt Leu					403
											gat Asp					451
-	-		-	-	-		_				gtg Val			_	_	499
-	-	-	_	•	_					-	ctt Leu 145	-	-			547
											ttt Phe					595
											cgt Arg					643
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											gca Ala					739
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											gtt Val					835
											cct Pro					883
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Lys Glu Ile Leu Ala Glu Phe Glu Ser

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Ala Asp Leu Gly Ala Thr Arg Leu Ala Ile Ile Thr Ala Pro Asn Lys 35 40 45

Asp Gly Ile Leu Lys His Phe Glu Glu Phe Pro Glu Leu Glu Ala Thr 50 55 60

Leu Glu Ala Arg Gly Lys Thr Asp Gln Leu Asn Lys Val Arg Ala Ala 65 70 75 80

Arg Glu Leu Ile Ala Thr Val Pro Val Val Gln Glu Lys Pro Leu Gly
85 90 95

Leu Gly His Ala Val Gly Leu Ala Glu Ser Val Leu Asp Asp Glu
100 105 110

Asp Val Val Ala Val Met Leu Pro Asp Asp Leu Val Leu Pro Phe Gly
115 120 125

Val Thr Glu Arg Met Ala Glu Val Arg Ala Lys Phe Gly Gly Ser Val 130 135 140

Leu Ala Ala Ile Glu Val Ala Glu Asp Glu Val Ser Asn Tyr Gly Val 145 150 155 160

Phe Lys Leu Gly Glu Leu Asp Ala Glu Ser Glu Ser Glu Gly Ile Arg 165 170 175

Arg Val Val Gly Met Val Glu Lys Pro Ala Pro Glu Asp Ala Pro Ser 180 185 190

Arg Phe Ala Ala Thr Gly Arg Tyr Leu Leu Asp Arg Ala Ile Phe Asp 195 200 205

Ala Leu Arg Arg Ile Glu Pro Gly Ala Gly Gly Glu Leu Gln Leu Thr 210 215 220

Asp Ala Ile Ala Leu Leu Ile Glu Glu Gly His Pro Val His Ile Val 225 230 235 240

Val His Glu Gly Lys Arg His Asp Leu Gly Asn Pro Ala Gly Tyr Ile

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Val Pro Lys Glu Leu Leu Pro Val Val Asp Thr Pro Gly Ile Glu Leu

Ile Ala Ala Glu Ala Ala Glu Leu Gly Ala Thr Arg Leu Ala Ile Ile

Thr Ala Pro Asn Lys Ala Gly Val Leu Ala His Phe Glu Arg Ser Ser 75

Glu Leu Glu Glu Thr Leu Met Glu Arg Gly Lys Thr Asp Gln Val Glu

Ile Ile Arg Arg Ala Ala Asp Leu Ile Lys Ala Val Pro Val Thr Gln 100 105 110

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163 gca ggg ctc gga tat gtt ggg ctt tca aat gca gct ctc ctc tct aaa Ala Gly Leu Gly Tyr Val Gly Leu Ser Asn Ala Ala Leu Leu Ser Lys

aat cat aaa gtt gtt gca gtt gac att gat gaa gaa cga gtg aaa cta 211 Asn His Lys Val Val Ala Val Asp Ile Asp Glu Glu Arg Val Lys Leu 25

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gct tcg Ala Ser 135															547
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ctg ggg Leu Gly		Glu													643
cct ccg Pro Pro	Ile I			_	-				_						691
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gat act Asp Thr 215															787
ggg gta Gly Val 230			_		-										835
ttt gga Phe Gly		Gly													883
gcc aac Ala Asn	Tyr I	_	-	-	_	_					_	-			931

gca aat aag act Ala Asn Lys Thr 280			Ile Al						979
tca cct act gta Ser Pro Thr Val 295				u Val M					1027
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Val Phe Gly His Val Asp Ala Ser Tyr Trp Arg Asp Met Gly Thr P 215 220 225	ro
agc gac ttc gtc cgc ggc tcg gct gac ctg gtc cgc ggc att gcg t Ser Asp Phe Val Arg Gly Ser Ala Asp Leu Val Arg Gly Ile Ala T 230 235 240 2	ac 835 yr 45
tcc cca ttg ctc gaa ggc aaa aca gga gag tcg ctt gtc gac gcc t Ser Pro Leu Leu Glu Gly Lys Thr Gly Glu Ser Leu Val Asp Ala S 250 255 260	
gcc ggc gtt cgc gac ggc gtc ctg ctc ggc gga acc gta gtc g Ala Gly Val Arg Asp Gly Val Leu Leu Leu Gly Gly Thr Val Val G 265 270 275	
cgc ggc act gag atc ggt gcc ggc tgc cgc gtt gac aac act gtt a Arg Gly Thr Glu Ile Gly Ala Gly Cys Arg Val Asp Asn Thr Val I 280 285 290	
ttc gac ggc gtc acc att gaa cca ggt gcg gtc att gaa aat tcc a Phe Asp Gly Val Thr Ile Glu Pro Gly Ala Val Ile Glu Asn Ser I 295 300 305	
att tcc tcg gga gca cgc atc ggt gct aat gcg cac atc tcc ggt t Ile Ser Ser Gly Ala Arg Ile Gly Ala Asn Ala His Ile Ser Gly C 310 315 320 3	
atc att ggc gag ggc gca cag gtt ggt gct cgg tgt gaa ctc aac g Ile Ile Gly Glu Gly Ala Gln Val Gly Ala Arg Cys Glu Leu Asn A 330 335 340	
ggg atg cgc gtc ttc cca ggc gtt gtg atc cca gac agc gga att c Gly Met Arg Val Phe Pro Gly Val Val Ile Pro Asp Ser Gly Ile A 345 350 355	
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<pre>&lt;400&gt; 274 Met Thr Leu Thr Asp Asn Ser Lys Asn Val Asp Ala Val Ile Leu V 1 5 10 15</pre>	al
Gly Gly Lys Gly Thr Arg Leu Arg Pro Leu Thr Val Asn Thr Pro L	ys
20 25 30	
	la
20 25 30  Pro Met Leu Pro Thr Ala Gly His Pro Phe Leu Thr His Leu Leu A	

Leu Glu Ile Glu Tyr Val Val Glu Asp Gln Pro Leu Gly Thr Gly Gly
85 90 95

Gly Ile Arg Asn Val Tyr Asp Lys Leu Arg His Asp Thr Ala Ile Val 100 105 110

Phe Asn Gly Asp Val Leu Ser Gly Ala Asp Leu Asn Ser Ile Leu Asp 115 120 125

Thr His Arg Glu Lys Asp Ala Asp Leu Thr Met His Leu Val Arg Val 130 135 140

Ala Asn Pro Arg Ala Phe Gly Cys Val Pro Thr Asp Glu Asp Gly Arg 145 150 155 160

Val Ser Glu Phe Leu Glu Lys Thr Glu Asp Pro Pro Thr Asp Gln Ile 165 170 175

Asn Ala Gly Cys Tyr Val Phe Lys Lys Glu Leu Ile Glu Gln Ile Pro 180 185 190

Ala Gly Arg Ala Val Ser Val Glu Arg Glu Thr Phe Pro Gln Leu Leu 195 200 205

Glu Glu Gly Lys Arg Val Phe Gly His Val Asp Ala Ser Tyr Trp Arg 210 215 220

Asp Met Gly Thr Pro Ser Asp Phe Val Arg Gly Ser Ala Asp Leu Val 225 230 235 240

Arg Gly Ile Ala Tyr Ser Pro Leu Leu Glu Gly Lys Thr Gly Glu Ser
245 250 255

Leu Val Asp Ala Ser Ala Gly Val Arg Asp Gly Val Leu Leu Gly 260 265 270

Gly Thr Val Val Gly Arg Gly Thr Glu Ile Gly Ala Gly Cys Arg Val 275 280 285

Asp Asn Thr Val Ile Phe Asp Gly Val Thr Ile Glu Pro Gly Ala Val 290 295 300

Ile Glu Asn Ser Ile Ile Ser Ser Gly Ala Arg Ile Gly Ala Asn Ala 305 310 315 320

His Ile Ser Gly Cys Ile Ile Gly Glu Gly Ala Gln Val Gly Ala Arg 325 330 335

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cag go Gln Al 21															787
ggc gg Gly G: 230															835
tat ga Tyr As															883
ggc ta Gly T	ac tg yr Tr <sub>l</sub>	g cgc p Arg 265	gac Asp	gtc Val	ggt Gly	acc Thr	att Ile 270	gat Asp	gcg Ala	ttc Phe	tac Tyr	gag Glu 275	tgc Cys	cac His	931
atg ga Met As		u Ile													979
tgg co Trp Pi 29															1027
gtt co Val A: 310															1075
att to															1123
gtc ga Val G															1171
cgc at Arg I		y Lys													1219
gtg g Val Va 3	tt gte al Va 75	c cgc l Arg	gac Asp	gga Gly	gag Glu 380	ctc Leu	atc Ile	ggt Gly	gtc Val	gac Asp 385	caa Gln	gtg Val	cgc Arg	gat Asp	1267
gcg ca Ala G 390															1315
aac ca Asn G		_	taaa	acgg	gaa a	aggga	acctt	ta aa	aa	٠					1350

<sup>&</sup>lt;210> 276

<sup>&</sup>lt;211> 409

<sup>&</sup>lt;212> PRT

<sup>&</sup>lt;213> Corynebacterium glutamicum

<sup>&</sup>lt;400> 276

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Ala	Gly	Gly	Glu 20	Gly	Lys	Arg	Leu	Phe 25	Pro	Leu	Thr	Glu	Asp 30	Arg	Ala
Lys	Pro	Ala 35	Val	Pro	Phe	Gly	Gly 40	Thr	Tyr	Arg	Leu	Ile 45	Asp	Phe	Val
Leu	Ser 50	Asn	Leu	Val	Asn	Ser 55	Gly	Phe	Leu	Lys	Ile 60	Ala	Val	Leu	Thr
Gln 65	Tyr	Lys	Ser	His	Ser 70	Leu	Asp	Arg	His	Ile 75	Ser	Leu	Ser	Trp	Asn 80
Val	Ser	Gly	Pro	Thr 85	Gly	Gln	Tyr	Ile	Ala 90	Ser	Val	Pro	Ala	Gln 95	Gln
Arg	Leu	Gly	Lys 100	Arg	Trp	Phe	Thr	Gly 105	Ser	Ala	Asp	Ala	Ile 110	Leu	Gln
Ser	Leu	Asn 115	Leu	Ile	Ser	Asp	Glu 120	Lys	Pro	Asp	Tyr	Val 125	Ile	Val	Phe
Gly	Ala 130	Asp	His	Val	Tyr	Arg 135	Met	Asp	Pro	Ser	Gln 140	Met	Leu	Asp	Glu
His 145	Ile	Ala	Ser	Gly	Arg 150	Ala	Val	Ser	Val	Ala 155	Gly	Ile	Arg	Val	Pro 160
Arg	Glu	Glu	Ala	Thr 165	Ala	Phe	Gly	Суѕ	Ile 170	Gln	Ser	Asp	Val	Asp 175	Gly
Asn	Ile	Thr	Glu 180	Phe	Leu	Glu	Lys	Pro 185	Ala	Asp	Pro	Pro	Gly 190	Thr	Pro
Asp	Asp	Pro 195	Asp	Met	Thr	Tyr	Ala 200	Ser	Met	Gly	Asn	Tyr 205	Ile	Phe	Thr
Thr	Glu 210	Ala	Leu	Ile	Gln	Ala 215	Leu	Lys	Asp	Asp	Glu 220	Asn	Asn	Glu	Asn
Ser 225	Asp	His	Asp	Met	Gly 230	Gly	Asp	Ile	Ile	Pro 235	Tyr	Phe	Val	Ser	Arg 240
Asn	Asp	Ala	His	Val 245	Tyr	Asp	Phe	Ser	Gly 250	Asn	Ile	Val	Pro	Gly 255	Ala
Thr	Glu	Arg	Asp 260	Lys	Gly	Tyr	Trp	Arg 265	Asp	Val	Gly	Thr	11e 270	Asp	Ala
Phe	Tyr	Glu 275	Cys	His	Met	Asp	Leu 280	Ile	Ser	Val	His	Pro 285	Ile	Phe	Asn
Leu	Tyr 290	Asn	Ser	Glu	Trp	Pro 295	Ile	His	Thr	Thr	Ser 300	Glu	Gly	Asn	Leu
Pro 305	Pro	Ala	Lys	Phe	Val 310	Arg	Gly	Gly	Ile	Ala 315	Gln	Ser	Ser	Met	Val 320
Ser	Ser	Gly	Ser	Ile	Ile	Ser	Ala	Gly	Thr	Val	Arg	Asn	Ser	Val	Leu

330

335

325

Ser Asn Asn Val Val Glu Glu Gly Ala Thr Val Glu Gly Ala Val 345 Leu Met Pro Gly Val Arg Ile Gly Lys Gly Ala Val Val Arg His Ala Ile Leu Asp Lys Asn Val Val Val Arg Asp Gly Glu Leu Ile Gly Val Asp Gln Val Arg Asp Ala Gln Arg Phe Lys Val Ser Ala Gly Gly Val Val Val Gly Lys Asn Gln Val Val 405 <210> 277 <211> 903 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(880) <223> RXN00014 <400> 277 catcaaagtg accgccggcg gcgtcgaatg gtccgttgca ggaaacgcgg aagcagttag 60 tgagatetee gaaactttaa gegeactaga etaacaacae atg age aaa tat gea Met Ser Lys Tyr Ala gac gat tta gcc tta gcc ctc gaa ctt gcc gaa ctt gcc gat tcc atc 163 Asp Asp Leu Ala Leu Ala Leu Glu Leu Ala Glu Leu Ala Asp Ser Ile 10 211 acc ctc gac cgc ttc gaa gcc tct gac ctg gaa gta tcc tcc aag cca Thr Leu Asp Arg Phe Glu Ala Ser Asp Leu Glu Val Ser Ser Lys Pro 259 gac atg act ccc gtc agc gat gcc gac ctg gcg acc gaa gaa gca ctc Asp Met Thr Pro Val Ser Asp Ala Asp Leu Ala Thr Glu Glu Ala Leu 45 307 cqt gag aaa atc gcc acc gcc cgc ccc gcc gac tcc atc ctc ggt gaa Arg Glu Lys Ile Ala Thr Ala Arg Pro Ala Asp Ser Ile Leu Gly Glu 60 gaa ttc ggt ggc gac gta gaa ttc agc ggc cgc cag tgg atc atc gac 355 Glu Phe Gly Gly Asp Val Glu Phe Ser Gly Arg Gln Trp Ile Ile Asp 403 ccc atc gac ggc acc aaa aac tac gtc cgc ggc gtc ccc gta tgg gca Pro Ile Asp Gly Thr Lys Asn Tyr Val Arg Gly Val Pro Val Trp Ala 451 acc ctg atc gcg ctg ctc gac aac ggc aaa ccc gtc gca ggt gtc atc

Thr Leu Ile Ala Leu Leu Asp Asn Gly Lys Pro Val Ala Gly Val Ile

105 110 115 tcc gca ccc gca ctg gct agg cgt tgg tgg gca tcc gaa ggg gcc ggc 499 Ser Ala Pro Ala Leu Ala Arg Arg Trp Trp Ala Ser Glu Gly Ala Gly gca tgg cgc acc ttc aac ggc agc tcc cca cgc aaa ctg tcc gtg tcc 547 Ala Trp Arg Thr Phe Asn Gly Ser Ser Pro Arg Lys Leu Ser Val Ser cag gtg tcc aag ctt gac gac gcc tcc ctc tcc tcc tcc ctc tcc 595 Gln Val Ser Lys Leu Asp Asp`Ala Ser Leu Ser Phe Ser Ser Leu Ser ggc tgg gcc gaa cga gat ttg cgc gat cag ttc gtc tcc cta act gat Gly Trp Ala Glu Arg Asp Leu Arg Asp Gln Phe Val Ser Leu Thr Asp 170 691 ace ace tog ega etc ege gge tac gge gae tte tte tec tac tge etc Thr Thr Trp Arg Leu Arg Gly Tyr Gly Asp Phe Phe Ser Tyr Cys Leu 190 gtc gcc gaa ggt gcc gtc gat atc gcc gct gaa cca gaa gtc agc ctc 739 Val Ala Glu Gly Ala Val Asp Ile Ala Ala Glu Pro Glu Val Ser Leu 200 205 tgg gat ctt gct ccc ctg tcc atc ctg gtc acc gaa gcc gga gga aag 787 Trp Asp Leu Ala Pro Leu Ser Ile Leu Val Thr Glu Ala Gly Gly Lys 215 tto acc toa ctg got ggo gto gat gga coa cac ggt ggo gat gca gta 835 Phe Thr Ser Leu Ala Gly Val Asp Gly Pro His Gly Gly Asp Ala Val 230 235 gcc acc aac ggc atc ctg cac gat gag acg ctg gat cgt tta aaa 880 Ala Thr Asn Gly Ile Leu His Asp Glu Thr Leu Asp Arg Leu Lys 250 255 260 903 tagactcccq qqttttqctt ggt <210> 278 <211> 260 <212> PRT <213> Corynebacterium glutamicum <400> 278 Met Ser Lys Tyr Ala Asp Asp Leu Ala Leu Glu Leu Ala Glu Leu Ala Asp Ser Ile Thr Leu Asp Arg Phe Glu Ala Ser Asp Leu Glu Val Ser Ser Lys Pro Asp Met Thr Pro Val Ser Asp Ala Asp Leu Ala 40 Thr Glu Glu Ala Leu Arg Glu Lys Ile Ala Thr Ala Arg Pro Ala Asp

75

Ser Ile Leu Gly Glu Glu Phe Gly Gly Asp Val Glu Phe Ser Gly Arg

Gln	Trp	Ile	Ile	Asp 85	Pro	Ile	Asp	Gly	Thr 90	Lys	Asn	Tyr	Val	Arg 95	Gly	
Val	Pro	Val	Trp 100	Ala	Thr	Leu	Ile	Ala 105	Leu	Leu	Asp	Asn	Gly 110	Lys	Pro	
Val	Ala	Gly 115	Val	Ile	Ser	Ala	Pro 120	Ala	Leu	Ala	Arg	Arg 125	Trp	Trp	Ala	
Ser	Glu 130	Gly	Ala	Gly	Ala	Trp 135	Arg	Thr	Phe	Asn	Gly 140	Ser	Ser	Pro	Arg	
Lys 145	Leu	Ser	Val	Ser	Gln 150	Val	Ser	Lys	Leu	Asp 155	Asp	Ala	Ser	Leu	Ser 160	
Phe	Ser	Ser	Leu	Ser 165	Gly	Trp	Ala	Glu	Arg 170	Asp	Leu	Arg	Asp	Gln 175	Phe	
Val	Ser	Leu	Thr 180	Asp	Thr	Thr	Trp	Arg 185	Leu	Arg	Gly	Tyr	Gly 190	Asp	Phe	
Phe	Ser	Tyr 195	Cys	Leu	Val	Ala	Glu 200	Gly	Ala	Val	Asp	Ile 205	Ala	Ala	Glu	
Pro	Glu 210	Val	Ser	Leu	Trp	Asp 215	Leu	Ala	Pro	Leu	Ser 220	Ile	Leu	Val	Thr	
Glu 225	Ala	Gly	Gly	Lys	Phe 230	Thr	Ser	Leu	Ala	Gly 235	Val	Asp	Gly	Pro	His 240	
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Asp	Arg	Leu	Lys 260													
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tgaç	gate	tcc (	gaaa	cttta	aa go	egcad	ctaga	a cta	agcaa	acac	-	-	aaa Lys		•	115
													gat Asp			163
													tcc Ser			211

25 30 35

						gat Asp										259
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						gaa Glu										355
						aac Asn										403
						gac Asp										451
tcc Ser	gca Ala	ccc Pro 120	gca Ala	ctg Leu	gct Ala	agg Arg	cgt Arg 125	tgg Trp	tgg Trp	gca Ala	tcc Ser	gaa Glu 130	ggg Gly	gcc Ala	ggc Gly	499
gca Ala	tgg Trp 135	cgc Arg	acc Thr	ttc Phe	aac Asn	ggc Gly 140	agc Ser	tcc Ser	cca Pro	cgc Arg	aaa Lys 145	ctg Leu	tcc Ser	gtg Val	tcc Ser	547
						gac Asp										595
						ttg Leu										643
acc Thr	acc Thr	tgg Trp	cga Arg 185	ctc Leu	cgc Arg	ggc Gly	tac Tyr	ggc Gly 190	gac Asp	ttc Phe	ttc Phe	tcc Ser	tac Tyr 195	tgc Cys	ctc Leu	691
						gat Asp										739
						tcc Ser 220										787
						gtc Val										835
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<212> PRT

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<400> 280

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Val Ser Ser Lys Pro Asp Met Thr Pro Val Ser Asp Ala Asp Leu Ala 35 40 45

Thr Glu Glu Ala Leu Arg Glu Lys Ile Ala Thr Ala Arg Pro Ala Asp 50 55 60

Ser Ile Leu Gly Glu Glu Phe Gly Gly Asp Val Glu Phe Ser Gly Arg
65 70 75 80

Gln Trp Ile Ile Asp Pro Ile Asp Gly Thr Lys Asn Tyr Val Arg Gly
85 90 95

Val Pro Val Trp Ala Thr Leu Ile Ala Leu Leu Asp Asn Gly Lys Pro 100 105 110

Val Ala Gly Val Ile Ser Ala Pro Ala Leu Ala Arg Arg Trp Trp Ala 115 120 125

Ser Glu Gly Ala Gly Ala Trp Arg Thr Phe Asn Gly Ser Ser Pro Arg 130 135 140

Lys Leu Ser Val Ser Gln Val Ser Lys Leu Asp Asp Ala Ser Leu Ser 145 150 155 160

Phe Ser Ser Leu Ser Gly Trp Ala Glu Arg Asp Leu Arg Asp Gln Phe 165 170 175

Val Ser Leu Thr Asp Thr Thr Trp Arg Leu Arg Gly Tyr Gly Asp Phe 180 185 190

Phe Ser Tyr Cys Leu Val Ala Glu Gly Ala Val Asp Ile Ala Ala Glu 195 200 205

Pro Glu Val Ser Leu Trp Asp Leu Ala Pro Leu Ser Ile Leu Val Thr 210 215 220

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Asp Arg Leu Lys 260

<210> 281

<211> 978

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ate que aaq tea ate aag eet tee teg egt gge gaa etg gaa ate ace Ile Ala Lys Ser Ile Lys Pro Ser Ser Arg Gly Glu Leu Glu Ile Thr

190

185

691

Ser Val Asn Asp Ala Tyr Leu Gln Gln Gly Ala Leu Thr Val Gln Arg 200 205 210	739
ctg gac cgt ggc gat gtc tgg tta gat acc ggc aca atc gat tcc atg Leu Asp Arg Gly Asp Val Trp Leu Asp Thr Gly Thr Ile Asp Ser Met 215 220 225	787
tcc gag gcg tct tcc tat gtt gag gtc ctg caa aaa cgt acc ggc aac Ser Glu Ala Ser Ser Tyr Val Glu Val Leu Gln Lys Arg Thr Gly Asn 230 235 240 245	835
atc atc gga tcc ccc gaa gtc gct gcg tac cgc gaa ggt ttc atc aca Ile Ile Gly Ser Pro Glu Val Ala Ala Tyr Arg Glu Gly Phe Ile Thr 250 255 260	883
gct gaa gaa ctc aca gtg ctt ggt gag gaa ctg aag aaa tca ggc tac Ala Glu Glu Leu Thr Val Leu Gly Glu Glu Leu Lys Lys Ser Gly Tyr 265 270 275	931
gga aac tac ctg ctg aga gct ttg taatttacgg tgtggttgtg gag Gly Asn Tyr Leu Leu Arg Ala Leu 280 285	978
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Val Lys Gly Ile Ile Leu Ala Gly Gly Ser Gly Thr Arg Leu Tyr Pro  1 5 10 15  Ile Thr Lys Gly Ile Ser Lys Gln Leu Met Pro Ile Tyr Asp Lys Pro	
Val Lys Gly Ile Ile Leu Ala Gly Gly Ser Gly Thr Arg Leu Tyr Pro 1 5 10 15  Ile Thr Lys Gly Ile Ser Lys Gln Leu Met Pro Ile Tyr Asp Lys Pro 20 25 30  Met Val Tyr Tyr Pro Leu Thr Thr Leu Ile Gln Ala Gly Ile Lys Asp	
Val Lys Gly Ile Ile Leu Ala Gly Gly Ser Gly Thr Arg Leu Tyr Pro 1 5 10 10 15 Pro 1 15 Pro 1 10 Thr Lys Gly Ile Ser Lys Gln Leu Met Pro Ile Tyr Asp Lys Pro 20 25 30  Met Val Tyr Tyr Pro Leu Thr Thr Leu Ile Gln Ala Gly Ile Lys Asp 35 40 40 Ser Ala Ser Phe Glu Arg Leu	
Val Lys Gly Ile Ile Leu Ala Gly Gly Ser Gly Thr Arg Leu Tyr Pro 1	
Val Lys Gly Ile Ile Leu Ala Gly Gly Ser Gly Thr Arg Leu Tyr Pro 10 15    Ile Thr Lys Gly Ile Ser Lys Gln Leu Met Pro Ile Tyr Asp Lys Pro 20    Met Val Tyr Tyr Pro Leu Thr Thr Leu Ile Gln Ala Gly Ile Lys Asp 35    Ile Leu Ile Ile Thr Thr Pro Glu Asp Ser Ala Ser Phe Glu Arg Leu 50    Leu Gly Asp Gly Ser Ser Trp Gly Ile Asn Leu Thr Tyr Ala Val Gln 80    Pro Ser Pro Asp Gly Leu Ala Gln Ala Phe Ile Ile Gly Glu Glu Phe	
Val Lys Gly Ile Ile Leu Ala Gly Gly Ser Gly Thr Arg Leu Tyr Pro 10 15 Ile Thr Lys Gly Ile Ser Lys Gln Leu Met Pro Ile Tyr Asp Lys Pro 20 Met Val Tyr Tyr Pro Leu Thr Thr Leu Ile Gln Ala Gly Ile Lys Asp 35 Ile Leu Ile Ile Thr Thr Pro Glu Asp Ser Ala Ser Phe Glu Arg Leu 60 Fro Ser Pro Asp Gly Ser Ser Trp Gly Ile Asn Leu Thr Tyr Ala Val Gln 80 Pro Ser Pro Asp Gly Leu Ala Gln Ala Phe Ile Ile Gly Glu Glu Phe 95 Ile Gly Asp Asp Asp Val Ala Leu Val Leu Gly Asp Asn Ile Phe Asp	
Val Lys Gly Ile Ile Leu Ala Gly Gly Ser Gly Thr Arg Leu Tyr Pro 10 15    Ile Thr Lys Gly Ile Ser Lys Gln Leu Met Pro Ile Tyr Asp Lys Pro 30    Met Val Tyr Tyr Pro Leu Thr Thr Leu Ile Gln Ala Gly Ile Lys Asp 45    Ile Leu Ile Ile Thr Thr Pro Glu Asp Ser Ala Ser Phe Glu Arg Leu 50    Leu Gly Asp Gly Ser Ser Trp Gly Ile Asn Leu Thr Tyr Ala Val Gln 80    Pro Ser Pro Asp Gly Leu Ala Gln Ala Phe Ile Ile Gly Glu Glu Phe 95    Ile Gly Asp Asp Asp Asp Val Ala Leu Val Leu Gly Asp Asn Ile Phe Asp 100    Gly Ala Gln Leu Gly His Ala Leu Lys Gln Cys Ser Asn Pro Asp Gly	

Thr Ala Pro Lys Ser Asn Phe Ala Val Val Gly Leu Tyr Phe Tyr Asp Asn Arg Val Val Asp Ile Ala Lys Ser Ile Lys Pro Ser Ser Arg Gly Glu Leu Glu Ile Thr Ser Val Asn Asp Ala Tyr Leu Gln Gln Gly Ala 200 Leu Thr Val Gln Arg Leu Asp Arg Gly Asp Val Trp Leu Asp Thr Gly Thr Ile Asp Ser Met Ser Glu Ala Ser Ser Tyr Val Glu Val Leu Gln 235 Lys Arg Thr Gly Asn Ile Ile Gly Ser Pro Glu Val Ala Ala Tyr Arg Glu Gly Phe Ile Thr Ala Glu Glu Leu Thr Val Leu Gly Glu Glu Leu 260 Lys Lys Ser Gly Tyr Gly Asn Tyr Leu Leu Arg Ala Leu 280 <210> 283 <211> 891 <212> DNA <213> Corynebacterium glutamicum <221> CDS <222> (101)..(868) <223> RXA02666 <400> 283 gctcggcgac gaggaagaga agaaggacgc attcgacgac ttcgacgatt ccgacgtgga 60 tottgacgat ctgagetteg acgacgaaga ttagacgeec atg teg tet aca ega Met Ser Ser Thr Arg 163 ate eee gte ate gea ete ete geg geg geg ggg ege gga ace ege ete Ile Pro Val Ile Ala Leu Leu Ala Ala Gly Arg Gly Thr Arg Leu 10 ggc gga ccc atc ccc aaa gca ttc gtc acg ttg cgt gaa cgc aca ctt 211 Gly Gly Pro Ile Pro Lys Ala Phe Val Thr Leu Arg Glu Arg Thr Leu 259 tta gag cgc tcg ctc caa gcc atg ctc acc tcc gaa agc gtc gac gaa Leu Glu Arg Ser Leu Gln Ala Met Leu Thr Ser Glu Ser Val Asp Glu ate ate etc gtc age ecc gac atg gaa ace tac gec egc gat ttg 307 Ile Ile Ile Leu Val Ser Pro Asp Met Glu Thr Tyr Ala Arg Asp Leu ctg cgc aaa cgc ggt ctt ttg aac gac ccc gaa ggg gta cgc gta cgg Leu Arg Lys Arg Gly Leu Leu Asn Asp Pro Glu Gly Val Arg Val Arg

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		a Thr Ala Val	atc cca gta ctg o Ile Pro Val Leu I 145	
			ggc gga gta gtt g Gly Gly Val Val V	
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		r Asp Asp Ala	agc ttg atg gaa t Ser Leu Met Glu 1 210	
		l Gln Gly Asp	cca atg gcg ttt a Pro Met Ala Phe I 225	
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	ctc tcc gat Leu Ser Asp				
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	gtt att ttc Val Ile Phe 75				
	gat cct ctt Asp Pro Leu 90				
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	at gat sp Asp														931
	ca ctc la Leu 280	Ser													979
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					ctg Leu											643
					ggc Gly											691
					cca Pro											739
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Gin Leu Leu Arg Ser vai Thr Arg Asp Pro Gly Pro Phe Int Ala Asp 145 150 155 160

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Asp Phe Asp Ala Leu Cys Tyr Leu Asn Pro Gly Ala Thr Pro Val Glu 180 185 190

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	ctt Leu															259
																307
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100

90

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PCT/IB00/00943 WO 01/00844

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414

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1 5 10 15  Glu Asp Met Ile Lys Thr Ile Thr Lys Thr Phe Val Ile Ala His Asp 25  Gln Asp Ser Asp Glu His Leu Ala Gln Ala Leu Val Tyr Asn Ala Gly 45  Arg Leu Ala Trp Arg Met Arg Glu Asn Gly Val Asp Thr Asp Tyr Lys 50  Thr Ser Val Ser Asp Val Val Thr Asp Ala Asp Arg Ala Ala Glu Ala	
Glu Asp Met Ile Lys Thr Ile Thr Lys Thr Phe Val Ile Ala His Asp 25 Thr Asp Ser Asp Glu His Leu Ala Gln Ala Leu Val Tyr Asn Ala Gly Asg Leu Ala Trp Arg Met Arg Glu Asn Gly Val Asp Thr Asp Tyr Lys 50 Thr Ser Val Ser Asp Val Val Thr Asp Ala Asp Arg Ala Ala Glu Ala 65 To 70 Thr Asp Arg Pro Glu Asp Gly Val Leu Phe Val Ala Gly Val Leu Glu Ala Leu Arg Pro Glu Asp Gly Val Leu	
Glu Asp Met Ile Lys Thr Ile Thr Lys Thr Phe Val Ile Ala His Asp 25 Thr Ser Val Asp Asp Met Arg Glu Asp Thr Asp Tyr Lys 50 Thr Ser Val Ser Asp Val Val Thr Asp Ala Asp Arg Ala Ala Glu Ala 65 To 70 Thr Asp Ala Asp Arg Ala Ala Glu Ala 65 Glu Glu Glu Glu Ala Asp Arg Ala Asp Arg Ala Ser Lys Ser Gly Lys Thr Trp Val	
Glu Asp Met Ile Lys Thr Ile Thr Lys Thr Phe Val Ile Ala His Asp 20 Ser Asp Glu His Leu Ala Gln Ala Leu Val Tyr Asn Ala Gly Arg Leu Ala Trp Arg Met Arg Glu Asn Gly Val Asp Thr Asp Tyr Lys 50 Feb Val Ala Gly Val Asp Thr Asp Tyr Lys 60 Feb Val Ala Gly Val Ala Gly Val Ala Glu Ala Glu Ala Gly Val Ala Gly Val Ala Gly Val Ala Glu Ala Glu Ala Gly Val Ala Ala Gly Val Ala Ala Gly Val Ala Ala Gly Val Ala Ala Gly Gly Glu Gly Ala Asp Arg Ala Ser Lys Ser Gly Lys Thr Trp Val Ilo Ala Ala Pro Val Ala Asp Gly Thr Tyr Asn Phe Thr Gln Gly Ser Asp Tyr	

Phe Gly Gly Pro Gly Ile Arg Thr Thr Leu Asp Gly Lys Glu Leu Asp Leu Leu Val Asp Ala Pro Leu Asn Gln Ile Ser Leu Ala Thr Tyr Ile 185 His Pro Ser Arq Ile Ala Glu Pro Asp Ile Gln Lys Ala Trp Met Ser 200 Val Ala Thr His Pro Ala Thr Leu Arg Met Phe Gly Ala Gly Ser Ile Asp Leu Ala Asn Ile Ala Asp Gly Ser Met Gly Ala Trp Val Gln His 230 235 Ser Val Ala Asp Trp Asp Trp Leu Pro Gly Arg Ala Leu Ile Glu Gly Val Gly Gly Ala Cys Ile Lys Val Thr Ala Gly Gly Val Glu Trp Ser Val Ala Gly Asn Ala Glu Ala Val Ser Glu Ile Ser Glu Thr Leu Ser Ala Leu Asp 290 <210> 293 <211> 948 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(925) <223> RXA01099 <400> 293 ggatgagggc attgattccg tcatcattgg caaggcactt tatgagcaca agttcaccct 60 cgaagaggct ttggctgcag tagaaaagct cggttaatac atg gat gct cgt ggg Met Asp Ala Arg Gly 163 atq ttg gcc att gcg gag gcc gtt gta gat gcc gaa gcc ctc ttc Met Leu Ala Ile Ala Glu Ala Val Val Asp Asp Ala Glu Ala Leu Phe 10 211 atg cag ggc ttc gga gct gca cct gcc cat atg aaa tcc ccg ggg gat Met Gln Gly Phe Gly Ala Ala Pro Ala His Met Lys Ser Pro Gly Asp 25 259 ttt gcc acg gaa gtg gat atg gcc atc gaa tcc cat atg cgt tcg atg Phe Ala Thr Glu Val Asp Met Ala Ile Glu Ser His Met Arg Ser Met 40 307 ctq aac atq atq aca qqc att qct qtc atc qqt gaa gaa ggt ggc ggt Leu Asn Met Met Thr Gly Ile Ala Val Ile Gly Glu Glu Gly Gly Gly 55 60

													ggc Gly			355
aac Asn	ttc Phe	gcg Ala	gcg Ala	tcc Ser 90	aac Asn	ccg Pro	atg Met	agc Ser	gcg Ala 95	atc Ile	ctg Leu	gtg Val	tct Ser	ttg Leu 100	ctt Leu	403
													atg Met 115			451
													aac Asn			499
													cac His			547
ttt Phe 150	agt Ser	tcc Ser	atg Met	gcc Ala	tcc Ser 155	ccg Pro	cgc Arg	aat Asn	aca Thr	gcg Ala 160	ttt Phe	cct Pro	gtg Val	gag Glu	ttg Leu 165	595
													cgt Arg			643
													cag Gln 195			691
		-	-	-	_		-			-		-	aat Asn		-	739
													gac Asp			787
	His	Pro	Trp	Ala		Gly	Arg	Gly	Val	Val	Ala		aca Thr			835
gct Ala	cac His	gat Asp	gtg Val	ctg Leu 250	tta Leu	agt Ser	aag Lys	att Ile	gaa Glu 255	aaa Lys	gtt Val	cgg Arg	ttg Leu	atg Met 260	cat His	883
					gac Asp											925
taaa	aatgg	ggc (	gtggd	caatt	c ga	ag										948

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<sup>&</sup>lt;211> 275

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Glu Tyr Lys 275

<210> 295

<211> 576

<212> DNA

<213> Corynebacterium glutamicum

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gca gaa gac gcg ccg ggt gca cag gcc ttc act cgc att gaa gat gct
Ala Glu Asp Ala Pro Gly Ala Gln Ala Phe Thr Arg Ile Glu Asp Ala
atc gca gcc gat gct gtc gac gca gtg ctg atc gcc gta cca ggt cag
Ile Ala Ala Asp Ala Val Asp Ala Val Leu Ile Ala Val Pro Gly Gln
ttc cat gag cca gta ctt gtc cca gca cta gaa gca ggc ctt ccc atc
                                                                   192
Phe His Glu Pro Val Leu Val Pro Ala Leu Glu Ala Gly Leu Pro Ile
     50
                         55
ctg tgt gaa aag cca ctg acc cca gat tct gaa tcc tca ctg cgc atc
                                                                   240
Leu Cys Glu Lys Pro Leu Thr Pro Asp Ser Glu Ser Ser Leu Arg Ile
65
                     70
gtc gag ctg gag cag aag ctg gac aag cca cac atc cag gtt ggt ttc
                                                                   288
Val Glu Leu Glu Gln Lys Leu Asp Lys Pro His Ile Gln Val Gly Phe
                 85
                                                                   336
atg cgc cgc ttc gac cct gag tac aac aac ttg cgc aaa ttg gtg gaa
Met Arg Arg Phe Asp Pro Glu Tyr Asn Asn Leu Arg Lys Leu Val Glu
            100
tee gge gaa get gge gaa etg ete atg ete ege gge etg eac ege aac
                                                                   384
Ser Gly Glu Ala Gly Glu Leu Leu Met Leu Arg Gly Leu His Arg Asn
        115
cca aqt qtt qqt qaq aqc tac acc cag tcc atg ctg atc acc gac tcc
                                                                   432
Pro Ser Val Gly Glu Ser Tyr Thr Gln Ser Met Leu Ile Thr Asp Ser
    130
                        135
                                                                   480
gto gto cao gaa tto gat gto ato coa tgg cto goa ggo too cga gtt
Val Val His Glu Phe Asp Val Ile Pro Trp Leu Ala Gly Ser Arg Val
145
                    150
gtc tcc gtt gaa gtg aag tac cca aag acc tcc tca ctg gcg cac tcc
                                                                   528
Val Ser Val Glu Val Lys Tyr Pro Lys Thr Ser Ser Leu Ala His Ser
ggc ctc aag gaa cca atc ctg gtg atc atg gag ctc gaa aac ggc gtg
Gly Leu Lys Glu Pro Ile Leu Val Ile Met Glu Leu Glu Asn Gly Val
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<sup>&</sup>lt;210> 296

<sup>&</sup>lt;211> 192

<sup>&</sup>lt;212> PRT

<sup>&</sup>lt;213> Corynebacterium glutamicum

<sup>&</sup>lt;400> 296

His Ile Ser Ala Ile Ile Glu Pro Asp Ala Ala Arg Ala Ala Ala Ala Ala Glu Asp Ala Pro Gly Ala Gln Ala Phe Thr Arg Ile Glu Asp Ala Ile Ala Ala Asp Ala Val Asp Ala Val Leu Ile Ala Val Pro Gly Gln Phe His Glu Pro Val Leu Val Pro Ala Leu Glu Ala Gly Leu Pro Ile Leu Cys Glu Lys Pro Leu Thr Pro Asp Ser Glu Ser Ser Leu Arg Ile Val Glu Leu Glu Gln Lys Leu Asp Lys Pro His Ile Gln Val Gly Phe Met Arg Arg Phe Asp Pro Glu Tyr Asn Asn Leu Arg Lys Leu Val Glu Ser Gly Glu Ala Gly Glu Leu Leu Met Leu Arg Gly Leu His Arg Asn 120 Pro Ser Val Gly Glu Ser Tyr Thr Gln Ser Met Leu Ile Thr Asp Ser 135 Val Val His Glu Phe Asp Val Ile Pro Trp Leu Ala Gly Ser Arg Val 150 . 155 Val Ser Val Glu Val Lys Tyr Pro Lys Thr Ser Ser Leu Ala His Ser 170 Gly Leu Lys Glu Pro Ile Leu Val Ile Met Glu Leu Glu Asn Gly Val 180

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ctg atc gcc gta cca ggt cag ttc cat gag cca gta ctt gtc cca gca Leu Ile Ala Val Pro Gly Gln Phe His Glu Pro Val Leu Val Pro Ala 48

96

35 40 45 192 cta gaa qca qqc ctt ccc atc ctg tgt gaa aag cca ctg acc cca gat Leu Glu Ala Gly Leu Pro Ile Leu Cys Glu Lys Pro Leu Thr Pro Asp 50 55 tot gaa too toa otg ogo ato gto gag otg gag cag aag otg gac aag 240 Ser Glu Ser Ser Leu Arg Ile Val Glu Leu Glu Gln Lys Leu Asp Lys 288 cca cac atc cag gtt ggt ttc atg cgc cgc ttc gac cct gag tac aac Pro His Ile Gln Val Gly Phe Met Arg Arg Phe Asp Pro Glu Tyr Asn 336 aac ttg cgc aaa ttg gtg gaa tcc ggc gaa gct ggc gaa ctg ctc atg Asn Leu Arg Lys Leu Val Glu Ser Gly Glu Ala Gly Glu Leu Leu Met 100 ctc cqc qqc ctq cac cqc aac cca agt gtt ggt gag agc tac acc cag 384 Leu Arg Gly Leu His Arg Asn Pro Ser Val Gly Glu Ser Tyr Thr Gln 115 tcc atg ctg atc acc gac tcc gtc gtc cac gaa ttc gat gtc atc cca Ser Met Leu Ile Thr Asp Ser Val Val His Glu Phe Asp Val Ile Pro 135 130 tgq ctc gca ggc tcc cga gtt gtc tcc gtt gaa gtg aag tac cca aag 480 Trp Leu Ala Gly Ser Arg Val Val Ser Val Glu Val Lys Tyr Pro Lys 145 150 ace tee tea etg geg cae tee gge etc aag gaa eea ate etg gtg ate Thr Ser Ser Leu Ala His Ser Gly Leu Lys Glu Pro Ile Leu Val Ile 170 175 165 549 atg gag ctc gaa aac ggc gtg Met Glu Leu Glu Asn Gly Val 180 <210> 298 <211> 183 <212> PRT <213> Corynebacterium glutamicum Ala Ala Arq Ala Ala Ala Ala Glu Asp Ala Pro Gly Ala Gln Ala Phe Thr Arg Ile Glu Asp Ala Ile Ala Ala Asp Ala Val Asp Ala Val Leu Ile Ala Val Pro Gly Gln Phe His Glu Pro Val Leu Val Pro Ala Leu Glu Ala Gly Leu Pro Ile Leu Cys Glu Lys Pro Leu Thr Pro Asp

Ser Glu Ser Ser Leu Arg Ile Val Glu Leu Glu Gln Lys Leu Asp Lys

Pro His Ile Gln Val Gly Phe Met Arg Arg Phe Asp Pro Glu Tyr Asn

90 85 95 Asn Leu Arg Lys Leu Val Glu Ser Gly Glu Ala Gly Glu Leu Leu Met 105 Leu Arg Gly Leu His Arg Asn Pro Ser Val Gly Glu Ser Tyr Thr Gln 120 Ser Met Leu Ile Thr Asp Ser Val Val His Glu Phe Asp Val Ile Pro 135 Trp Leu Ala Gly Ser Arg Val Val Ser Val Glu Val Lys Tyr Pro Lys 150 155 Thr Ser Ser Leu Ala His Ser Gly Leu Lys Glu Pro Ile Leu Val Ile 170 Met Glu Leu Glu Asn Gly Val 180 <210> 299 <211> 1128 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(1105) <223> RXA01632 <400> 299 aagggetgea acgtgettte gacaccacca tegeagegtt tgaacaaget getegteteg 60 cccctccac taactgatct ttgaaaggct gaaaaaactc atg act ctt cgt atc Met Thr Leu Arg Ile gcc ctt ttc ggc gct ggc cgc atc ggt cac gtc cac gct gcc aac att Ala Leu Phe Gly Ala Gly Arg Ile Gly His Val His Ala Ala Asn Ile 10 gct gca aac cct gat ctt gaa ctc gtt gtt atc gcc gat cct ttc att 211 Ala Ala Asn Pro Asp Leu Glu Leu Val Val Ile Ala Asp Pro Phe Ile 25 qaa qqc qca cag cqt ttq qca qaa qcc aat qqq qca gaa gcg gtt gca 259 Glu Gly Ala Gln Arg Leu Ala Glu Ala Asn Gly Ala Glu Ala Val Ala 40 45 tca cca gat gag gtg ttc gcc cgc gat gat atc gat ggc atc gtg atc 307 Ser Pro Asp Glu Val Phe Ala Arg Asp Asp Ile Asp Gly Ile Val Ile 55 60 355 ggt tea eea ace age ace eac gtt gat etg ate ace ege gee gtg gaa Gly Ser Pro Thr Ser Thr His Val Asp Leu Ile Thr Arg Ala Val Glu 70 75 403 cgt ggc att cct gca ctg tgc gaa aaa ccc att gat tta gac att gaa Arg Gly Ile Pro Ala Leu Cys Glu Lys Pro Ile Asp Leu Asp Ile Glu

			gac ggc gct tcc aag gtg Asp Gly Ala Ser Lys Val 115	451
			tct ttc gct gcc atc aat Ser Phe Ala Ala Ile Asn 130	499
gcg cga gtg gca Ala Arg Val Ala 135	aac cag gag Asn Gln Glu 140	atc ggc aac Ile Gly Asn	ctg gag cag ttg gtg atc Leu Glu Gln Leu Val Ile 145	547
			gac tac atc gca ggt tcc Asp Tyr Ile Ala Gly Ser 160 165	595
,, ,,			gat ctg gat atg gcg cgt Asp Leu Asp Met Ala Arg 180	643
			gca acc ggc gcc aat gtt Ala Thr Gly Ala Asn Val 195	691
			tac gac cag gtt atc gtc Tyr Asp Gln Val Ile Val 210	739
			aac atc gtg aac tcc cgc Asn Ile Val Asn Ser Arg 225	787
			gag gct ttc ggc tct aag Glu Ala Phe Gly Ser Lys 240 245	835
	•		acc acg gtg cgc aag cac Thr Thr Val Arg Lys His 260	883
			att ttc aac ttc ttc ctc Ile Phe Asn Phe Phe Leu 275	931
			ctc gca act ttt gct caa Leu Ala Thr Phe Ala Gln 290	979
			aac ttc gag gac ggc gtc Asn Phe Glu Asp Gly Val 305	1027
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cct 1128

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<211> 335

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<213> Corynebacterium glutamicum

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His Ala Ala Asn Ile Ala Ala Asn Pro Asp Leu Glu Leu Val Val Ile 20 25 30

Ala Asp Pro Phe Ile Glu Gly Ala Gln Arg Leu Ala Glu Ala Asn Gly 35 40 45

Ala Glu Ala Val Ala Ser Pro Asp Glu Val Phe Ala Arg Asp Asp Ile 50 55 60

Asp Gly Ile Val Ile Gly Ser Pro Thr Ser Thr His Val Asp Leu Ile 65 70 75 80

Thr Arg Ala Val Glu Arg Gly Ile Pro Ala Leu Cys Glu Lys Pro Ile 85 90 95

Asp Leu Asp Ile Glu Met Val Arg Ala Cys Lys Glu Lys Ile Gly Asp 100 105 110

Gly Ala Ser Lys Val Met Leu Gly Phe Asn Arg Arg Phe Asp Pro Ser 115 120 125

Phe Ala Ala Ile Asn Ala Arg Val Ala Asn Gln Glu Ile Gly Asn Leu 130 135 140

Glu Gln Leu Val Ile Ile Ser Arg Asp Pro Ala Pro Ala Pro Lys Asp 145 150 155 160

Tyr Ile Ala Gly Ser Gly Gly Ile Phe Arg Asp Met Thr Ile His Asp 165 170 175

Leu Asp Met Ala Arg Phe Phe Val Pro Asn Ile Val Glu Val Thr Ala 180 185 190

Thr Gly Ala Asn Val Phe Ser Gln Glu Ile Ala Glu Phe Asn Asp Tyr 195 200 205

Asp Gln Val Ile Val Thr Leu Arg Gly Ser Lys Gly Glu Leu Ile Asn 210 215 220

Ile Val Asn Ser Arg His Cys Ser Tyr Gly Tyr Asp Gln Arg Leu Glu 225 230 235 240

Ala Phe Gly Ser Lys Gly Met Leu Ala Ala Asp Asn Ile Arg Pro Thr 245 250 255

Thr Val Arg Lys His Asn Ala Glu Ser Thr Glu Gln Ala Asp Pro Ile 260 265 270

Phe Asn Phe Phe Leu Glu Arg Tyr Asp Ala Ala Tyr Lys Ala Glu Leu

285

280

275

120

Ala Thr Phe Ala Gln Gly Ile Arg Asp Gly Gln Gly Phe Ser Pro Asn Phe Glu Asp Gly Val Ile Ala Leu Glu Leu Ala Asn Ala Cys Leu Glu 315 . Ser Ala Gln Thr Gly Arg Thr Val Thr Leu Asn Pro Ala Asn Val <210> 301 <211> 1206 <212> DNA <213> Corvnebacterium glutamicum <221> CDS <222> (101)..(1183) <223> RXA01633 <400> 301 qcqaatqcat qccttqaatc aqctcaaacc gqccqcaccq tcaccctcaa ccctgccaac 60 gtttagtcaa cgtctagtta atgcctaagg agaaaacctc atg aaa aac atc acc Met Lys Asn Ile Thr atc gga atg gtc ggc gtc ggc cgc att ggc cgc atg cac gtc gcc aac 163 Ile Gly Met Val Gly Val Gly Arg Ile Gly Arg Met His Val Ala Asn atg ctt gcc gtt gct gaa act ttg aag gaa cgc gac ctc aac att gag 211 Met Leu Ala Val Ala Glu Thr Leu Lys Glu Arg Asp Leu Asn Ile Glu ate gtg etc gea gac gea atg eec ggt ttt geg gag eag gtg gge geg 259 Ile Val Leu Ala Asp Ala Met Pro Gly Phe Ala Glu Gln Val Gly Ala 45 307 gac atg ggc gtg aag gcg gcg gca agc gtc gat aag ctt att gag gac Asp Met Gly Val Lys Ala Ala Ala Ser Val Asp Lys Leu Ile Glu Asp 355 ggg gtg gat gcc ctt ttc att gcc acc agc acc gct ggc cac gtc gat Gly Val Asp Ala Leu Phe Ile Ala Thr Ser Thr Ala Gly His Val Asp gtt ttg cgc aag ggc atc gcg gca aag ctg ccg atg ttc tgc gag aag 403 Val Leu Arg Lys Gly Ile Ala Ala Lys Leu Pro Met Phe Cys Glu Lys 451 ccg atc gcg tcg gat gtg cct gag tcg ctg aac atc atc cgc gaa att Pro Ile Ala Ser Asp Val Pro Glu Ser Leu Asn Ile Ile Arg Glu Ile 110 499 gat gcg gct ggc gcg acg gtt cag gtc ggc cac cag cgc cgt ttt gac Asp Ala Ala Gly Ala Thr Val Gln Val Gly His Gln Arg Arg Phe Asp

ctc ggt tac ca Leu Gly Tyr Gl 135	g gaa gct aaa Glu Ala Lys 140	cga cgc cta Arg Arg Leu	gat gca ggc Asp Ala Gly 145	gac ctc Asp Leu	ggc 547 Gly
tgg ctt cat tc Trp Leu His Se 150					
gtg tcc tac tg Val Ser Tyr Cy			Phe Arg Asp		
cac gat ttc ga His Asp Phe As 18	lle Ile Arg				
gtg tac gcc aa Val Tyr Ala Ly 200					
ggt gac atc ga Gly Asp Ile As 215	acc gga gcg Thr Gly Ala 220	Ala Leu Leu	acg ctt gcc Thr Leu Ala 225	gac ggc Asp Gly	acc 787 Thr
ctc gcc acc gc Leu Ala Thr Al 230					
cgc ctc gat gt Arg Leu Asp Va			Thr Ile Val		
gaa aag tct gc Glu Lys Ser Al 26	a Phe Ala Ser				
ggc gaa tcg ca Gly Glu Ser Hi 280					
aat gag tgc at Asn Glu Cys Il 295		Glu Leu Ile			
cct tgt acc cc Pro Cys Thr Pr 310					
gct cag ctg tc Ala Gln Leu Se			Val Lys Ile		
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<211> 361

<212> PRT

<213> Corynebacterium glutamicum

<400> 302

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Met His Val Ala Asn Met Leu Ala Val Ala Glu Thr Leu Lys Glu Arg  $20 \hspace{1cm} 25 \hspace{1cm} 30$ 

Asp Leu Asn Ile Glu Ile Val Leu Ala Asp Ala Met Pro Gly Phe Ala 35 40 45

Glu Gln Val Gly Ala Asp Met Gly Val Lys Ala Ala Ala Ser Val Asp 50 55 60

Lys Leu Ile Glu Asp Gly Val Asp Ala Leu Phe Ile Ala Thr Ser Thr 65 70 75 80

Ala Gly His Val Asp Val Leu Arg Lys Gly Ile Ala Ala Lys Leu Pro 85 90 95

Met Phe Cys Glu Lys Pro Ile Ala Ser Asp Val Pro Glu Ser Leu Asn 100 105 110

Ile Ile Arg Glu Ile Asp Ala Ala Gly Ala Thr Val Gln Val Gly His
115 120 125

Gln Arg Arg Phe Asp Leu Gly Tyr Gln Glu Ala Lys Arg Arg Leu Asp 130 135 140

Ala Gly Asp Leu Gly Trp Leu His Ser Leu Lys Ala Val Ser Ser Asp 145 150 155 160

Ala Phe Pro Pro Pro Val Ser Tyr Cys Ala Thr Ser Gly Gly Leu Phe 165 170 175

Arg Asp Val Ser Leu His Asp Phe Asp Ile Ile Arg Trp Leu Thr Gly
180 185 190

Gln Asp Ile Val Glu Val Tyr Ala Lys Gly Ser Asn Asn Gly Asp Pro 195 200 205

Glu Ile Gly Ala Val Gly Asp Ile Asp Thr Gly Ala Ala Leu Leu Thr 210 215 220

Leu Ala Asp Gly Thr Leu Ala Thr Ala Ile Ala Thr Arg Tyr Asn Gly 225 230 235 240

Ala Gly His Asp Val Arg Leu Asp Val Met Gly Ser Lys Asp Ser Thr 245 250 255

Ile Val Gly Leu Asp Glu Lys Ser Ala Phe Ala Ser Ala Glu Glu Gly 260 265 270

Ile Asp Phe Pro Thr Gly Glu Ser His Pro Thr Phe Ala Glu Arg Phe 275 280 285

Ala Asp Ala Tyr Lys Asn Glu Cys Ile Ala Phe Val Glu Leu Ile Leu

290 295 300 Gly Glu Arg Glu Asn Pro Cys Thr Pro Ala Asp Ala Val Ala Ala Ala 310 Ile Val Ala Asp Ala Ala Gln Leu Ser Leu Val Thr Gly Glu Pro Val Lys Ile Pro Thr Val Arg Glu Ile Leu Glu Gly Ser Ala Gln Pro Val Glu Val Arg Ala Leu Val Pro Ser Ala 360 <210> 303 <211> 1146 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(1123) <223> RXN01406 <400> 303 gttcctcatt cctctaatcg gcgcactatc tttgcctcgc gacggcggtg cccgagcctt 60 ttcctcctct tagaaaccca cttctgaaag gtataaaaac atg act att cga atc Met Thr Ile Arq Ile 1 qqa ctc qtt qqc tac qqt gtc qqc qgc agg ctc ttt cac acc cct tac Gly Leu Val Gly Tyr Gly Val Gly Gly Arg Leu Phe His Thr Pro Tyr 10 atc caa gct tct acg cac tgc gaa tta gta ggc gta gtt gct cgt tcc 211 Ile Gln Ala Ser Thr His Cys Glu Leu Val Gly Val Val Ala Arg Ser 25 259 gaa ggc acc aaa gca gcc gtt gca gaa gat ctt cca gat gtt gcc atc Glu Gly Thr Lys Ala Ala Val Ala Glu Asp Leu Pro Asp Val Ala Ile 45 gtg gga tcg ctg aca gaa ctc ctc gaa ctg ggc gtc gat gca gtg gtg 307 Val Gly Ser Leu Thr Glu Leu Leu Glu Leu Gly Val Asp Ala Val Val 55 atc tcc acc cct cca gcc acg cgc cgg gaa ctg gcc ttg gaa gca atc 355 Ile Ser Thr Pro Pro Ala Thr Arg Arg Glu Leu Ala Leu Glu Ala Ile 70 aac gca ggt gtc gca gtg gtt gcc gat aaa ccg ttt gca cca tca gcc Asn Ala Gly Val Ala Val Val Ala Asp Lys Pro Phe Ala Pro Ser Ala gea gat gec atg gaa ett gte gaa gee gee gaa aag get gga gtg etg Ala Asp Ala Met Glu Leu Val Glu Ala Ala Glu Lys Ala Gly Val Leu

ctc aac gtc ttc cac aac agg cgc aac gac acc cac att gtc acg gca

Leu	Asn	Val 120	Phe	His	Asn	Arg	Arg 125	Asn	Asp	Thr	His	11e 130	Val	Thr	Ala	
ctg Leu	gga Gly 135	atc Ile	caa Gln	gaa Glu	gaa Glu	ctt Leu 140	ggt Gly	gcg Ala	atg Met	cgt Arg	gga Gly 145	ctg Leu	gac Asp	ctg Leu	cga Arg	547
	gac Asp															595
ttg Leu	ctg Leu	cgc Arg	gat Asp	ctg Leu 170	ggc Gly	tca Ser	cac His	gta Val	gtc Val 175	gat Asp	cag Gln	acc Thr	ctg Leu	gtt Val 180	ctc Leu	643
	ggg Gly															691
	gaa Glu															739
	ggc Gly 215															787
	tgg Trp															835
	acc Thr															883
aat Asn	gac Asp	cgc Arg	gaa Glu 265	cac His	tgg Trp	ggc Gly	tac Tyr	gaa Glu 270	tcg Ser	gag Glu	gag Glu	cgg Arg	tgg Trp 275	ggc Gly	acc Thr	931
	gtt Val															979
tac Tyr	acc Thr 295	cgc Arg	ttc Phe	tac Tyr	gat Asp	gcc Ala 300	ttt Phe	gcc Ala	ttg Leu	gct Ala	gtg Val 305	gaa Glu	aac Asn	ggt Gly	ggc Gly	1027
gca Ala 310	ggg Gly	ccg Pro	gtg Val	cct Pro	gca Ala 315	cgt Arg	gaa Glu	ggt Gly	gtt Val	gca Ala 320	gtg Val	ctc Leu	aag Lys	gtg Val	ttg Leu 325	1075
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<213> Corynebacterium glutamicum

<400> 304

Met Thr Ile Arg Ile Gly Leu Val Gly Tyr Gly Val Gly Gly Arg Leu 1 5 10 15

Phe His Thr Pro Tyr Ile Gln Ala Ser Thr His Cys Glu Leu Val Gly 20 25 30

Val Val Ala Arg Ser Glu Gly Thr Lys Ala Ala Val Ala Glu Asp Leu 35 40 45

Pro Asp Val Ala Ile Val Gly Ser Leu Thr Glu Leu Leu Glu Leu Gly 50 55 60

Val Asp Ala Val Val Ile Ser Thr Pro Pro Ala Thr Arg Arg Glu Leu 65 70 75 80

Ala Leu Glu Ala Ile Asn Ala Gly Val Ala Val Val Ala Asp Lys Pro 85 90 95

Phe Ala Pro Ser Ala Ala Asp Ala Met Glu Leu Val Glu Ala Ala Glu
100 105 110

Lys Ala Gly Val Leu Leu Asn Val Phe His Asn Arg Arg Asn Asp Thr 115 120 125

His Ile Val Thr Ala Leu Gly Ile Gln Glu Glu Leu Gly Ala Met Arg 130 135 140

Gly Leu Asp Leu Arg Leu Asp Leu Ile Glu Pro Asp Ser Leu Glu Ala 145 150 155 160

Gly Pro Glu Gly Gly Leu Leu Arg Asp Leu Gly Ser His Val Val Asp 165 170 175

Gln Thr Leu Val Leu Met Gly Pro Ala Thr Ser Val Thr Ala Gln Leu 180 185 190

Gly Ser Ile Asp Leu Pro Glu Gly Pro Thr Asn Ala Arg Phe Arg Ile 195 200 205

Val Leu Glu His Glu Ser Gly Ala Val Ser His Ile Ser Ala Ser Lys 210 215 220

Ile Asp Arg Leu Glu Ser Trp Glu Ile Arg Leu Val Gly Glu Arg Gly 225 230 235 240

Ser Tyr Val Ser Asn Tyr Thr Asp Val Gln Thr Val Ala Ile Lys Gln 245 250 255

Gly Leu Arg Pro Thr Asn Asp Arg Glu His Trp Gly Tyr Glu Ser Glu 260 265 270

Glu Arg Trp Gly Thr Leu Val Thr Asp Glu Gly Ser Lys Val Ile Pro 275 280 285

Ser Ala Gln Gly Asp Tyr Thr Arg Phe Tyr Asp Ala Phe Ala Leu Ala 290 295 300

Val Glu Asn Gly Gly Ala Gly Pro Val Pro Ala Arg Glu Gly Val Ala

310

305

315

320

Val Leu Lys Val Leu Asp Ala Val Ala Gln Ser Ala Ala Glu Lys Arg 330 325 Thr Ile Glu Leu Ser 340 <210> 305 <211> 1200 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(1177) <223> RXN01630 <400> 305 gtaggtgagt cttcgtgaga tacccccggc cagtcataca gttcaaccaa gctccaccac 60 ccagataaaa acctgcgggt tgcgttttag gagaattccc atg agt gat caa aaa 115 Met Ser Asp Gln Lys 1 att qtt qtt qqc ctq cta ggc atc acc cac ccg cat gcg tcg gcg cgg 163 Ile Val Val Gly Leu Leu Gly Ile Thr His Pro His Ala Ser Ala Arg 10 gtg cgt gcc ctc cgt gaa att gat ggg gta gag gtc gtc gcc gcc gcg 211 Val Arg Ala Leu Arg Glu Ile Asp Gly Val Glu Val Val Ala Ala Ala 25 30 gat act gat tcc cgc ctc cag tac ttc acc gac aaa tat gat gtt gaa 259 Asp Thr Asp Ser Arg Leu Gln Tyr Phe Thr Asp Lys Tyr Asp Val Glu 40 45 ccc cgc gag atc gat gac gtc ttg aac gac gat cgc atc aac gcc atc Pro Arg Glu Ile Asp Asp Val Leu Asn Asp Asp Arg Ile Asn Ala Ile 55 60 atg qtt cac tcc aag agc aag gac atg gtc cct cac gcc aag cgc gcg 355 Met Val His Ser Lys Ser Lys Asp Met Val Pro His Ala Lys Arg Ala 70 ctc qcq qcc qqa aaa tcc qtc qtc qtg gag aag ccc ggc ggg gga aca 403 Leu Ala Ala Gly Lys Ser Val Val Glu Lys Pro Gly Gly Gly Thr 100 90 gtg gcg gat ctt gag gag ctc ctg gcc ctc aaa gaa gct gcc gat cct 451 Val Ala Asp Leu Glu Glu Leu Leu Ala Leu Lys Glu Ala Ala Asp Pro 105 115 cag cga atc gtg cag gtc ggg tac aac gtc cgc ctg tct gaa tcg gtt 499 Gln Arg Ile Val Gln Val Gly Tyr Asn Val Arg Leu Ser Glu Ser Val 125 120 547 cag aga tta aaa gag ctt ctc gac gcc ggc ctc atc ggc gaa gtc gtc Gln Arg Leu Lys Glu Leu Leu Asp Ala Gly Leu Ile Gly Glu Val Val 135 140

					gcc Ala									595
				_	gac Asp	-			 _					643
					ttg Leu									691
	-	_			aag Lys		-			-	-		_	739
					gca Ala 220									787
					cac His									835
-	-	-			acc Thr	_		_	-	-				883
					tac Tyr									931
_	 				acc Thr	_	_				_		_	979
					ttc Phe 300									1027
-		_		_	atg Met	_						Arg		1075
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<213> Corynebacterium glutamicum

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Thr Pro Pro Phe Ala Arg Thr Glu Ser Asn Lys Phe Ser Glu Leu Pro

Glu Leu Glu Asn Ile Ser Asn Phe Arg Thr Glu Met Gln Gly Trp Val

433

295

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Leu	Thr	Val	Ala 340	Arg	Ile	Val	Ser	Ala 345	Суѕ	Tyr	Glu	Ser	Asp 350	Asn	Asn	
Gln	Gly	Ile 355	Ser	Val	Asn	Ile										
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tegg	jtege	gg (	ccgtt	atci	ct tt	taaq	gagga	a gaa	aatti	tag	atg Met 1	agc Ser	acg Thr	tcc Ser	acc Thr 5	115
atc Ile	agg Arg	gtt Val	gcc Ala	att Ile 10	gcc Ala	gga Gly	gtc Val	gga Gly	aac Asn 15	tgc Cys	gcg Ala	acc Thr	tcc Ser	ctc Leu 20	att . Ile	163
cag Gln	ggt Gly	gtg Val	gaa Glu 25	tat Tyr	tac Tyr	cga Arg	aat Asn	gcg Ala 30	gat Asp	cct Pro	tcc Ser	gaa Glu	act Thr 35	gtc Val	ccg Pro	211
ggt Gly	ttg Leu	atg Met 40	cac His	gtc Val	aaa Lys	ttc Phe	ggt Gly 45	gat Asp	tac Tyr	cac His	gtt Val	ggc Gly 50	gac Asp	att Ile	gaa Glu	259
		Ala		Phe	Asp	Val			gaa Glu							307
									tgc Cys							355
									cgt Arg 95							403
									gac Asp							451
									gca Ala							499

tcc tac Ser Tyr 135	Leu														547
gcc gcc Ala Ala 150															595
atc gcc Ile Ala	tcc Ser	gac Asp	cct Pro 170	gag Glu	tgg Trp	gct Ala	aag Lys	aag Lys 175	ttc Phe	act Thr	gac Asp	gct Ala	ggc Gly 180	atc Ile	643
cca att	gtt Val	ggc Gly 185	gat Asp	gac Asp	atc Ile	aaa Lys	tcc Ser 190	cag Gln	atc Ile	ggt Gly	gca Ala	acc Thr 195	atc Ile	acc Thr	691
cac cg		Leu													739
cgc acc Arg Th	r Met														787
ctt gad Leu Asy 230															835
gtg acc Val Th															883
cgc aad Arg Asi															931
cgc aa Arg Ly		Āla													979
ccc cty Pro Les 29	Asn	Leu	Glu	Tyr		Leu	Glu	Val	Trp	Asp	Ser				1027
gcc gg Ala Gl 310	c atc y Ile	atc Ile	atc Ile	gac Asp 315	gct Ala	gtt Val	cgc Arg	gcc Ala	gcc Ala 320	aag Lys	atc Ile	gcc Ala	ctc Leu	gat Asp 325	1075
cgc gg Arg Gl															1123
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<213> Corynebacterium glutamicum

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Ser Glu Thr Val Pro Gly Leu Met His Val Lys Phe Gly Asp Tyr His 35 40 45

Val Gly Asp Ile Glu Phe Val Ala Ala Phe Asp Val Asp Ala Glu Lys  $50 \hspace{1cm} 55 \hspace{1cm} 60$ 

Val Gly Ile Asp Leu Ala Asp Ala Thr Glu Ala Ser Gln Asn Cys Thr 65 70 75 80

Ile Lys Ile Ala Asp Val Pro Gln Thr Gly Ile Asn Val Leu Arg Gly 85 90 95

Pro Thr Leu Asp Gly Leu Gly Asp His Tyr Arg Ala Thr Ile Asp Glu 100 105 110

Ser Thr Ala Glu Pro Val Asp Val Val Gln Ala Leu Ile Asp Ala Lys 115 120 125

Ala Asp Val Leu Val Ser Tyr Leu Pro Val Gly Ser Glu Glu Ala Asp 130 135 140

Lys Phe Tyr Ala Gln Ala Ala Ile Asp Ala Gly Cys Ala Phe Val Asn 145 150 155 160

Ala Leu Pro Val Phe Ile Ala Ser Asp Pro Glu Trp Ala Lys Lys Phe 165 170 175

Thr Asp Ala Gly Ile Pro Ile Val Gly Asp Asp Ile Lys Ser Gln Ile 180 185 190

Gly Ala Thr Ile Thr His Arg Val Leu Ala Arg Leu Phe Glu Glu Arg 195 200 205

Gly Val Arg Val Asp Arg Thr Met Gln Leu Asn Val Gly Gly Asn Met 210 215 220

Asp Phe Lys Asn Met Leu Asp Arg Asn Arg Leu Glu Ser Lys Lys Val 225 230 235 240

Ser Lys Thr Gln Ala Val Thr Ser Asn Ile Pro Asp Gly Pro Leu Ser 245 250 255

Gly Lys Val Glu Asp Arg Asn Val His Ile Gly Pro Ser Asp His Val 260 265 270

Gln Trp Leu Asp Asp Arg Lys Trp Ala Tyr Val Arg Leu Glu Gly Thr 275 280 285

Ala Phe Gly Gly Val Pro Leu Asn Leu Glu Tyr Lys Leu Glu Val Trp

300 290 295 Asp Ser Pro Asn Ser Ala Gly Ile Ile Ile Asp Ala Val Arg Ala Ala 315 310 Lys Ile Ala Leu Asp Arg Gly Ile Gly Gly Pro Ile Met Pro Ala Ser 330 Ser Tyr Leu Met Lys Ser Pro Pro Glu Gln Leu Pro Asp Asp Val Ala Cys Glu Arg Leu Glu Ala Phe Ile Ile Glu Ala <210> 309 <211> 795 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(772) <223> RXN03057 <400> 309 catcaacgcc gagtacaact aaggacaact gataatgaca aatgctgcaa ttgtcggatg 60 aggagacgtc gcaaccgttc atacagaagc gctggaagct ttg gct tcc gat ctt 115 Leu Ala Ser Asp Leu ggt att aag ttc gtc gca gtg gtg gat aaa gat cta gag act gct gag Gly Ile Lys Phe Val Ala Val Val Asp Lys Asp Leu Glu Thr Ala Glu 10 aaa ttt gcg acg gga ctt gga gct gct ggc gat tct tca gaa agc agc 211 Lys Phe Ala Thr Gly Leu Gly Ala Ala Gly Asp Ser Ser Glu Ser Ser 25 259 gtc aag gcc cac ggc agc ctg ccg gct ttg ttc tcc aaa aag aag atc Val Lys Ala His Gly Ser Leu Pro Ala Leu Phe Ser Lys Lys Ile 40 45 307 gat gtt cta cac atc acc acc ccc cac gac caa cac att ggt ttg gct Asp Val Leu His Ile Thr Thr Pro His Asp Gln His Ile Gly Leu Ala 55 ctc gaa gcg cta cac cac ggt gta aat gtc atc ctg gaa aag ccg ttg 355 Leu Glu Ala Leu His His Gly Val Asn Val Ile Leu Glu Lys Pro Leu 70 gct aat gag ttg gac cag gcg cag cgt ctc atc gac tac ttg gat gaa 403 Ala Asn Glu Leu Asp Gln Ala Gln Arg Leu Ile Asp Tyr Leu Asp Glu aac ccc gat ggt cca aag att gca gtg tgc tat cag aac cgt tac aac 451 Asn Pro Asp Gly Pro Lys Ile Ala Val Cys Tyr Gln Asn Arg Tyr Asn 105

499

gtt tcc tcc cag gaa ctg cgt cgt ctg ctc gat tca ggt gac ctc ggt

Val Ser Ser 120		u Leu	Arg	Arg 125	Leu	Leu	Asp	Ser	Gly 130	Asp	Leu	Gly	
gcc atc aat Ala Ile Asn 135													547
tac tac acc Tyr Tyr Thr 150	cag aa Gln Ly	a cct s Pro 155	tgg Trp	cgt Arg	ggc Gly	cag Gln	caa Gln 160	gca Ala	cac His	tcc Ser	ggt Gly	ggt Gly 165	595
ggc ctg ctg Gly Leu Leu		n Gln											643
ttc ctt gga Phe Leu Gly													691
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aca													795
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<pre>&lt;211&gt; 224 &lt;212&gt; PRT &lt;213&gt; Coryn &lt;400&gt; 310 Leu Ala Ser 1 Leu Glu Thr Ser Ser Glu 35 Ser Lys Lys 50 His Ile Gly</pre>	Asp Le Ala Gl 20 Ser Se Lys Il Leu Al	u Gly  u Lys  r Val  e Asp  a Leu  70	Ile Phe Lys Val 55 Glu	Lys Ala Ala 40 Leu Ala	Phe Thr 25 His	10 Gly Gly Ile	Leu Ser Thr His 75	Gly Leu Thr 60 Gly	Ala Pro 45 Pro Val	Ala 30 Ala His	15 Gly Leu Asp Val	Asp Phe Gln Ile 80	
<pre>&lt;211&gt; 224 &lt;212&gt; PRT &lt;213&gt; Coryn &lt;400&gt; 310 Leu Ala Ser</pre>	Asp Le Ala Gl 20 Ser Se Lys Il Leu Al Pro Le 8	u Gly  Lys  r Val  e Asp  a Leu  70  u Ala	Ile Phe Lys Val 55 Glu Asn	Lys Ala Ala 40 Leu Ala Glu	Phe Thr 25 His Leu Leu	Gly Gly Ile His	Leu Ser Thr His 75 Gln	Gly Leu Thr 60 Gly	Pro 45 Pro Val	Ala 30 Ala His Asn	15 Gly Leu Asp Val Leu 95	Asp Phe Gln Ile 80 Ile	
<pre>&lt;211&gt; 224 &lt;212&gt; PRT &lt;213&gt; Coryn &lt;400&gt; 310 Leu Ala Ser 1 Leu Glu Thr  Ser Ser Glu 35 Ser Lys Lys 50 His Ile Gly 65 Leu Glu Lys</pre>	Asp Le Ala Gl 20 Ser Se Lys Il Leu Al Pro Le 8 Asp Gl 100 Tyr As	u Gly  Lys  r Val  e Asp  a Leu  70  u Ala  5	Ile Phe Lys Val 55 Glu Asn	Lys Ala Ala 40 Leu Ala Glu Asp	Phe Thr 25 His Leu Leu Gly 105	Gly Gly Ile His Asp 90 Pro	Leu Ser Thr His 75 Gln Lys	Gly Leu Thr 60 Gly Ala Ile	Ala Pro 45 Pro Val Gln Ala	Ala 30 Ala His Asn Arg Val	15 Gly Leu Asp Val Leu 95 Cys	Asp Phe Gln Ile 80 Ile	

140 130 135 Thr Arg Thr Pro Gly Tyr Tyr Thr Gln Lys Pro Trp Arg Gly Gln Gln 150 Ala His Ser Gly Gly Gly Leu Leu Met Asn Gln Ala Ile His Thr Leu Asp Leu Leu Gln Trp Phe Leu Gly Lys Ala Thr Glu Val Lys Gly Thr Val Ser Thr Asp Lys Tyr Ala Asp Val Ile Asp Val Glu Asp Thr Ala His Ala Tyr Ile Gly His Glu Ser Gly Val His Thr Ser Glu Val Ser <210> 311 <211> 795 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(772) <223> FRXA02902 <400> 311 catcaacgcc gagtacaact aaggacaact gataatgaca aatgctgcaa ttgtcggatg 60 aggagacgte geaacegtte atacagaage getggaaget ttg get tee gat ett Leu Ala Ser Asp Leu ggt att aag ttc gcc gca gtg gtg gat aaa gat cta gag act gct gag 163 Gly Ile Lys Phe Val Ala Val Val Asp Lys Asp Leu Glu Thr Ala Glu 10 15 aaa ttt gcg acg gga ctt gga gct gct ggc gat tct tca gaa agc agc 211 Lys Phe Ala Thr Gly Leu Gly Ala Ala Gly Asp Ser Ser Glu Ser Ser 25

gat gtt cta cac atc acc ccc cac gac caa cac att ggt ttg gct
Asp Val Leu His Ile Thr Thr Pro His Asp Gln His Ile Gly Leu Ala
55

ctc gaa gcg cta cac cac ggt gta aat gtc atc ctg gaa aag ccg ttg
Leu Glu Ala Leu His His Gly Val Asn Val Ile Leu Glu Lys Pro Leu
70

75

80

307

307

gtc aag gcc cac ggc agc ctg ccg gct ttg ttc tcc aaa aag aag atc Val Lys Ala His Gly Ser Leu Pro Ala Leu Phe Ser Lys Lys Ile

45

40

259

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gcc atc aat ggt gca tat tcc tct gtg gtg tgg acc cgc acc cca gg Ala Ile Asn Gly Ala Tyr Ser Ser Val Val Trp Thr Arg Thr Pro Gl 135	
Tyr Tyr Thr Gln Lys Pro Trp Arg Gly Gln Gln Ala His Ser Gly Gl 150 155 160	У
ggc ctg ctg atg aac caa gca att cac acc ctg gat ctg ctg cag tg Gly Leu Leu Met Asn Gln Ala Ile His Thr Leu Asp Leu Leu Gln Tr 170 175 180	
ttc ctt gga aag gca aca gaa gtc aag ggc act gtc tcc acc gat aag Phe Leu Gly Lys Ala Thr Glu Val Lys Gly Thr Val Ser Thr Asp Lys 185 190 195	
tat gcc gat gtc atc gat gtt gaa gac acc gca cac gca tac atc gg Tyr Ala Asp Val Ile Asp Val Glu Asp Thr Ala His Ala Tyr Ile Gl 200 205 210	
cac gag tcc gga gtc cac acc agt gaa gtg agt tgaaccatgc tattggt His Glu Ser Gly Val His Thr Ser Glu Val Ser 215 220	gat 792
aca	795
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105 110 100 Gln Asn Arg Tyr Asn Val Ser Ser Gln Glu Leu Arg Arg Leu Leu Asp 120 Ser Gly Asp Leu Gly Ala Ile Asn Gly Ala Tyr Ser Ser Val Val Trp 135 Thr Arg Thr Pro Gly Tyr Tyr Thr Gln Lys Pro Trp Arg Gly Gln Gln Ala His Ser Gly Gly Leu Leu Met Asn Gln Ala Ile His Thr Leu Asp Leu Leu Gln Trp Phe Leu Gly Lys Ala Thr Glu Val Lys Gly Thr 185 Val Ser Thr Asp Lys Tyr Ala Asp Val Ile Asp Val Glu Asp Thr Ala 200 His Ala Tyr Ile Gly His Glu Ser Gly Val His Thr Ser Glu Val Ser 215 220

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gcg gtc gtt acc gga gcg acc gga ggc atg gga att gag atc gtc aaa 163 Ala Val Val Thr Gly Ala Thr Gly Gly Met Gly Ile Glu Ile Val Lys 10 15 20

gac ctc tcc cgc gac cac att gtc tac gcc ttg ggc cga aat cca gag 211 Asp Leu Ser Arg Asp His Ile Val Tyr Ala Leu Gly Arg Asn Pro Glu 25 30 35

cat ctg gca gct ctc gca gag atc gag gga gta gag cct atc gag tcc 259 His Leu Ala Ala Leu Ala Glu Ile Glu Gly Val Glu Pro Ile Glu Ser 40 45 50

gat atc gtg aag gaa gtg ttg gaa gag gga ggc gtc gac aag cta aaa 307 Asp Ile Val Lys Glu Val Leu Glu Glu Gly Gly Val Asp Lys Leu Lys 55 60 65

aac ctc gac cac gtg gat acg ctg gtg cac gcc gcg gcg gtg gcg cgt 355

Asn Leu Asp His Val Asp Thr Leu Val His Ala Ala Ala Val Ala Arg 70 75 80 85	
gac acg acc atc gaa gcc ggc agt gtg gcc gaa tgg cac gca cac ctt Asp Thr Thr Ile Glu Ala Gly Ser Val Ala Glu Trp His Ala His Leu 90 95 100	403
gat ctc aac gtc att gtc ccg gcc gag ttg agt cgc caa ctc ttg ccc Asp Leu Asn Val Ile Val Pro Ala Glu Leu Ser Arg Gln Leu Leu Pro 105 110 115	451
gcc ctc cgc gcg gca tcc ggc tgc gtc atc tac atc aac tcc ggc gcc Ala Leu Arg Ala Ala Ser Gly Cys Val Ile Tyr Ile Asn Ser Gly Ala 120 125 130	499
ggc aac gga cca cac ccc ggc aac acc atc tac gcc gcc agc aaa cac Gly Asn Gly Pro His Pro Gly Asn Thr Ile Tyr Ala Ala Ser Lys His 135 140 145	547
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gct ggc gaa acc acc cag atc acc aac gtg gac gta cga cca cgt atc Ala Gly Glu Thr Thr Gln Ile Thr Asn Val Asp Val Arg Pro Arg Ile 215 220 225	787
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Gly Arg Asn Pro Glu His Leu Ala Ala Leu Ala Glu Ile Glu Gly Val 35 40 45	
Glu Pro Ile Glu Ser Asp Ile Val Lys Glu Val Leu Glu Glu Gly Gly 50 55 60	

Val Asp Lys Leu Lys Asn Leu Asp His Val Asp Thr Leu Val His Ala 65 Ala Ala Val Ala Arg Asp Thr Thr Ile Glu Ala Gly Ser Val Ala Glu Trp His Ala His Leu Asp Leu Asn Val Ile Val Pro Ala Glu Leu Ser 105 Arg Gln Leu Leu Pro Ala Leu Arg Ala Ala Ser Gly Cys Val Ile Tyr 120 Ile Asn Ser Gly Ala Gly Asn Gly Pro His Pro Gly Asn Thr Ile Tyr 135 140 Ala Ala Ser Lys His Ala Leu Arg Gly Leu Ala Asp Ala Phe Arg Lys Glu Glu Ala Asn Asn Gly Ile Arg Val Ser Thr Val Ser Pro Gly Pro 170 175 165 Thr Asn Thr Pro Met Leu Gln Gly Leu Met Asp Ser Gln Gly Thr Asn 185 Phe Arg Pro Glu Ile Tyr Ile Glu Pro Lys Glu Ile Ala Asn Ala Ile 200 195 Arg Phe Val Ile Asp Ala Gly Glu Thr Thr Gln Ile Thr Asn Val Asp 215 Val Arg Pro Arg Ile Glu Leu Ala Asp Arg Lys Asp 235 <210> 315 <211> 1008 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(985) <223> RXN02654 <400> 315 tattttcgga aatttataca gcaatcctcg aaatcctaat aaagatccct tatcgtqgga 60 gaggtacggt agttcgttcg aggacaacgt cgagaaaggc atg att tca ttg cta 115 Met Ile Ser Leu Leu aat gat cca cgt acg cta ttc ccg aaa gtc gat ccc cca aag caa agc Asn Asp Pro Arg Thr Leu Phe Pro Lys Val Asp Pro Pro Lys Gln Ser. 10 cag ccg gaa cca ggc cta gat ata aaa ctt tcc ccc caa gcc gat att 211 Gln Pro Glu Pro Gly Leu Asp Ile Lys Leu Ser Pro Gln Ala Asp Ile 25 ggt ctc tcc agc tat caa gga agt gga agg ctt aag ggc cgc aag gct 259

Gly	Leu	Ser 40	Ser	Tyr	Gln	Gly	Ser 45	Gly	Arg	Leu	Lys	Gly 50	Arg	Lys	Ala	
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	gct Ala															355
caa Gln	gcc Ala	gat Asp	gct Ala	gac Asp 90	aga Arg	gtg Val	ctc Leu	caa Gln	gca Ala 95	atc Ile	gag Glu	gaa Glu	aca Thr	ggt Gly 100	caa Gln	403
	gct Ala															451
	ctg Leu															499
	aac Asn 135															547
	gac Asp	_			_	-		_	_	_					-	595
	cgg Arg															643
	atc Ile			_			-					_	_			691
_	gat Asp		-	•		-	_	_	_			_		_		739
	gca Ala 215															787
	ggt Gly															835
	aaa Lys															883
	cct Pro															931
-	agc Ser			-		-		-		-				-		979

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1008

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Lys Gly Arg Lys Ala Leu Ile Thr Gly Gly Asp Ser Gly Ile Gly Ala

Ala Val Ala Ile Ala Tyr Ala Arg Glu Gly Ala Asp Val Ala Ile Ala

Tyr Leu Pro Glu Glu Gln Ala Asp Ala Asp Arg Val Leu Gln Ala Ile

Glu Glu Thr Gly Gln Lys Ala Phe Ser Phe Pro Gly Asp Leu Arg Asp

Pro Glu Tyr Cys Arg Ser Leu Val Gln Glu Thr Val Asn Ala Leu Gly

Gly Leu Asp Ile Leu Val Asn Asn Ala Ser Arg Gln Val Trp Ala Pro

Gly Leu Thr Glu Ile Thr Asp Glu Asn Phe Asp Gln Thr Leu Gln Val

Asn Leu Tyr Gly Ser Phe Arg Val Thr Lys Ala Ala Ile Pro His Leu

Lys Pro Gly Ser Ser Ile Ile Phe Thr Ser Ser Ile Gln Ala Tyr Gln

Pro Ser Glu Thr Leu Leu Asp Tyr Ala Met Thr Lys Ala Ala Leu Asn 195

Asn Leu Ser Lys Gly Leu Ala Ser Ser Leu Ile Gly Asp Gly Ile Arg 215

Val Asn Ser Val Ala Pro Gly Pro Phe Trp Thr Pro Leu Gln Pro Ser 225

His Gly Gln Pro Gln Glu Lys Ile Glu Gly Phe Gly Gln His Ala Pro 245 250

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gaa Glu	caa Gln 55	att Ile	cgt Arg	gat Asp	gcc Ala	att Ile 60	gct Ala	gtt Val	gcg Ala	gtg Val	gca Ala 65	cga Arg	gly ggg	tgg Trp	tct Ser	307
gtt Val 70	gtt Val	ggg Gly	cgc Arg	ggt Gly	gga Gly 75	gga Gly	agc Ser	tcg Ser	gtt Val	gct Ala 80	gga Gly	aat Asn	gcg Ala	atc Ile	ggt Gly 85	355
	ggt Gly															403 .
att Ile	gat Asp	cca Pro	gtt Val 105	gca Ala	caa Gln	act Thr	gca Ala	gtt Val 110	gtg Val	gaa Glu	ccc Pro	ggt Gly	gtg Val 115	gtg Val	tgt Cys	451
gat Asp	gcc Ala	ttg Leu 120	cgc Arg	gat Asp	gca Ala	gcc Ala	gca Ala 125	gaa Glu	ttc Phe	gga Gly	tta Leu	act Thr 130	tac Tyr	ggc Gly	ccg Pro	499
	cct Pro 135															547
	gcg Ala								Phe							595
	gtg Val															643
	aaa Lys															691
tta Leu	gcg Ala	tcc Ser 200	aag Lys	aat Asn	cag Gln	Asp	ctt Leu 205	att Ile	agt Ser	aaa Lys	Glu	ctg Leu 210	ggt Gly	cgt Arg	ttc Phe	739
	cgc Arg 215															787
	aaa Lys															835
_	acg Thr		_	_	-									-	•	883
-	gct Ala		-	_	-		-	_	-	_	-	_	-			931
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gct gcg ctg cgc Ala Ala Leu Arg 295		Gly Gln Ser			
cca gga aac cgc Pro Gly Asn Arg 310	atc ggc att Tle Gly Ile 315	gaa gcc ggc Glu Ala Gly	gga tgg ttg Gly Trp Leu 320	Tyr Cys	gag 1075 Glu 325
aca gga agt gac Thr Gly Ser Asp					
gca acc gcc gtt Ala Thr Ala Val 345	Asp Thr Ile				
gaa atg cgg gaa Glu Met Arg Glu 360					
acg cgc tta gct Thr Arg Leu Ala 375	, ,,,,,	Glu Ala Trp		-	-
gcg gtg cct cca Ala Val Pro Pro 390	• -	•		Leu Tyr	• •
ctg atg gat aag Leu Met Asp Lys	_				• •
gaa ggc tgc gtc Glu Gly Cys Val 425	His Val Arg		-	_	_
ggc ctg aag aaa Gly Leu Lys Lys 440		=		_	
gcg tot tat ggt Ala Ser Tyr Gly 455		Ser Gly Glu			
cgc tca tcc ttc Arg Ser Ser Phe 470		=		Arg Ala	
ttc gaa gaa ttc Phe Glu Glu Phe		-			-
gga gtg ttg gtc Gly Val Leu Val 505	Trp Ala Asp			-	-
ccg ggc cag cgc Pro Gly Gln Arg	•	_	•		

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gca tgc c Ala Cys A 550												1795
ggc gac g Gly Asp G												1843
atg ttc c Met Phe A												1891
aat gaa g Asn Glu A . 6		-	_	Ser	-		-	-			-	1939
tgt cca g Cys Pro V 615			-				-	-		-	-	1987
aaa cac t Lys His T 630	-		_		_	-			-	-		2035
tgg ctg c Trp Leu P	-		-	_				_				2083
acg ctt a Thr Leu I	-	-			_			_				2131
cgc aag g Arg Lys V 6			-	_	_	_				_		2179
cgc tcg c Arg Ser L 695												2227
gtg ctg t Val Leu T 710												2275
cac gca g His Ala A	la Ile											2323
cca gat g Pro Asp G												2371
ttg agc a Leu Ser M 7	_	-	-	_		-	-			_		2419

ccc tac ctg gad Pro Tyr Leu Asp 775								2467			
acc gtc atg ct Thr Val Met Lew 790								2515			
ctg gca cgc cti Leu Ala Arg Leu	gca gca Ala Ala 810	ctg acc Leu Thr	Lys P	cca ttc Pro Phe	gct gag Ala Glu	Val I	itc gca le Ala 20	2563			
cca aag atc acc Pro Lys Ile Thr 829	Glu Leu	gtc gag Val Glu	tct g Ser G 830	gga agc Gly Ser	ctc cag Leu Gln	cta a Leu T 835	ica gaa hr Glu	2611			
tca act gcg ct Ser Thr Ala Lev 840	acc cag Thr Gln	gtg cac Val His 845	tgc c Cys H	cac gag His Glu	cgt tcg Arg Ser 850	cta g Leu G	gc gac Sly Asp	2659			
cca caa caa tco Pro Gln Gln Sen 855								2707			
caa att gcc act Gln'Ile Ala Th 870	ggt tgt Gly Cys 875	tgc ggg Cys Gly	ctt g Leu A	gcc gga Ala Gly 880	aac tgg Asn Trp	ggc t Gly F	tt gaa Phe Glu 885	2755			
aaa gac cac gc Lys Asp His Ala			Ala I			Glu I		2803			
ccc aag gtc aga Pro Lys Val Arc 909	Lys Ala	gaa gga Glu Gly	cat g His V 910	gtg att /al Ile	gct gac Ala Asp	ggt t Gly F 915	tc tcc he Ser	2851			
tgc cgc acc cac Cys Arg Thr Gli 920								2899			
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Gln His Ser Arg	_	Leu Arg	Thr A	Asp Met	Thr Thr	Arg A	ala Ala				

Tyr Ser Ser Asp Ala Gly Ile Phe Arg Arg Val Pro Ala Ala Val Ala

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Gly	Asn	Ala	Ile	Gly 85	Glu	Gly	Leu	Ile	Ile 90	Asp	Thr	Ser	Arg	Tyr 95	Phe
Asn	Arg	Ile	Leu 100	Asp	Ile	Asp	Pro	Val 105	Ala	Gln	Thr	Ala	Val 110	Val	Glu
Pro	Gly	Val 115	Val	Суѕ	Asp	Ala	Leu 120	Arg	Asp	Ala	Ala	Ala 125	Glu	Phe	Gly
Leu	Thr 130	Tyr	Gly	Pro	Asp	Pro 135	Ser	Thr	His	Ser	Arg 140	Cys	Thr	Ile	Gly
Gly 145	Met	Val	Ala	Asn	Asn 150	Ala	Cys	Gly	Ser	His 155	Ser	Val	Ala	Phe	Gly 160
Thr	Ala	Ala	Glu	Asn 165	Leu	Val	Asp	Val	Thr 170	Leu	Met	Leu	Ser	Asp 175	Gly
Arg	Glu	Val	Thr 180	Val	Thr	Lys	Asp	Gly 185	Суѕ	Asp	Asp	Ala	Glu 190	Ile	Asn
Gln	Lys	Leu 195	Thr	Asp	Leu	Ala	Ser 200	Lys	Asn	Glņ	Asp	Leu 205	Ile	Ser	Lys
Glu	Leu 210	Gly	Arg	Phe	Pro	Arg 215	Gln	Val	Ser	Gly	Tyr 220	Gly	Leu	His	Tyr
Leu 225	Ala	His	Asp	Met	Ala 230	Lys	Ala	Met	Ala	Gly 235	Thr	Glu	Gly	Thr	Ile 240
Gly	Ile	Ile	Thr	Arg 245	Leu	Thr	Val	Lys	Leu 250	Val	Pro	Thr	Pro	Lys 255	Val
Lys	Ala	Leu	Ala 260	Val	Leu	Ala	Phe	Asp 265	Thr	Val	Phe	Asp	Ala 270	Ala	Arg
Ala	Ala	Ala 275	Lys	Leu	Arg	Leu	Pro 280	Gly	Val	Ala	Thr	11e 285	Glu	Gly	Met
Gly	Gly 290	Asp	Leu	Leu	Ala	Ala 295	Leu	Arg	Ser	Lys	Gln 300	Gly	Gln	Ser	Glu
Ala 305	Gly	Gln	Asn	Leu	Pro 310	Gly	Asn	Arg	Ile	Gly 315	Ile	Glu	Ala	Gly	Gly 320
Trp	Leu	Tyr	Cys	Glu 325	Thr	Gly	Ser	Asp	Thr 330	Leu	Gln	Ala	Ala	Val 335	Gln
Ala	Ala	Glu	Glu 340	Val	Ala	Thr	Ala	Val 345	Asp	Thr	Ile	Asp	Tyr 350	Val	Val
Val	Ser	Glu 355	Pro	Ser	Glu	Met	Arg 360	Glu	Leu	Trp	Arg	Ile 365	Arg	Glu	Ser

Ser Ala Gly Ile Val Thr Arg Leu Ala Asp Gly Glu Ala Trp Pro Asn Trp Glu Asp Ser Ala Val Pro Pro Glu Asn Leu Ala Asp Tyr Leu 390 Arg Asp Leu Tyr Ala Leu Met Asp Lys Phe Asp Tyr Gln Gly Ile Pro Phe Gly His Phe Gly Glu Gly Cys Val His Val Arg Ile Ser Phe Asp Phe Ser Thr Lys Glu Gly Leu Lys Lys Phe Glu Ala Phe Met Asn Glu Ala Ser Thr Leu Val Ala Ser Tyr Gly Gly Ser Leu Ser Gly Glu His Gly Asp Gly Arg Ala Arg Ser Ser Phe Leu Asp Arg Met Tyr Ser Ala Glu Met Arg Ala Leu Phe Glu Glu Phe Lys Leu Ile Phe Asp Pro Gln 490 Arg Ile Phe Asn Pro Gly Val Leu Val Trp Ala Asp Pro Val Met Gln Gly Leu Arg Met Asp Pro Gly Gln Arg Ala Leu Asp Ile Thr Pro Val His Lys Phe Ser Lys Asp Lys Gly Ser Met Ile Asn Ala Val Asn Arg Cys'Val Gly Val Ser Ala Cys Arg Ser Glu Ser Asp Ala Met Cys Pro Ser Phe Gln Ile Thr Gly Asp Glu Val His Ser Thr Arg Gly Arg Ala 570 Arg Leu Leu Ser Glu Met Phe Arg Gly Glu Ser Ile Ala Asp Gly Tyr Arg Ser Glu Glu Val Asn Glu Ala Leu Asp Leu Cys Leu Ser Cys Lys 600 Ala Cys Ala Ser Glu Cys Pro Val Asn Val Asp Met Ser Thr Tyr Lys Ala Glu Phe Leu Asp Lys His Tyr Ala Gly Arg Leu Arg Pro Met Ala 630 His Tyr Val Met Gly Trp Leu Pro Leu Leu Gly His Val Ala His Lys Ile Pro Leu Leu Pro Thr Leu Ile Asp Ala Thr Met Gln Ser Ala Leu Thr Ala Pro Val Val Arg Lys Val Gly Gly Leu Ala Asp Arg Pro Leu 675 680

Ile Ser Phe Ala His Arg Ser Leu Arg Lys Tyr Lys Pro Lys Lys Asn Ser Gly Glu Thr Val Val Leu Trp Pro Asp Ser Phe Asn Thr Asn Leu Asp Thr Gly Pro Ala His Ala Ala Ile Lys Thr Leu Glu Ala Leu Gly Tyr Asn Val Val Ile Pro Asp Gly Phe Val Cys Cys Gly Leu Thr Trp His Ser Thr Gly Gln Leu Ser Met Thr Lys Lys Val Leu Glu Gln Thr Ala Lys Val Met Lys Pro Tyr Leu Asp Gln Gly Leu Thr Val Val Gly Leu Glu Pro Ser Cys Thr Val Met Leu Gln Asp Glu Ala Thr Glu Leu Ser Asp Asn Pro Asp Leu Ala Arg Leu Ala Ala Leu Thr Lys Pro Phe Ala Glu Val Ile Ala Pro Lys Ile Thr Glu Leu Val Glu Ser Gly Ser Leu Gln Leu Thr Glu Ser Thr Ala Leu Thr Gln Val His Cys His Glu 840 Arg Ser Leu Gly Asp Pro Gln Gln Ser Ala Leu Val Leu Glu Ala Leu 855 Gly Val Lys Asp Glu Gln Ile Ala Thr Gly Cys Cys Gly Leu Ala Gly 870 875 Asn Trp Gly Phe Glu Lys Asp His Ala Glu Met Ser Phe Ala Leu Gly Glu Arg Glu Leu Phe Pro Lys Val Arg Lys Ala Glu Gly His Val Ile Ala Asp Gly Phe Ser Cys Arg Thr Gln Ile Glu Gln Gly Thr Gly Lys Gln Ala Thr His Leu Ala Glu Val Val Leu Ser Ile Leu Glu Gln Asn Asn Met Ala Gln 945

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cgt Arg	ttc Phe 210	cct Pro	cgc Arg	caa Gln	gtg Val	tcg Ser 215	ggc Gly	tac Tyr	ggt Gly	ttg Leu	cat His 220	tat Tyr	ctt Leu	gcc Ala	cac His	672
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				cct Pro												1200
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					ggt Gly											1776
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atg Met	ggc Gly	tgg Trp	Leu	ccg Pro 645	ctt Leu	Leu	gga Gly	His	Val	Ala	cat His	Lys	Ile	Pro	Leu	1968
ctt Leu	cct Pro	acg Thr	ctt Leu 660	atc Ile	gac Asp	gcc Ala	acc Thr	atg Met 665	cag Gln	tca Ser	gca Ala	ctc Leu	acc Thr 670	gcc Ala	cca Pro	2016
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					cgc Arg											2112
					ccc Pro 710											2160
cca	gct	cac	gca	gcg	atc	aaa	act	ctt	gaa	gcc	ctc	ggt	tac	aac	gtg	2208

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ggc caa tt Gly Gln Le 75	eu Ser Met	aca aag Thr Lys	aaa gto Lys Val 760	cta gaa Leu Glu	caa acg Gln Thr 765	Ala Lys	gtg 2304 Val
atg aaa co Met Lys Pr 770	cc tac ctg co Tyr Leu	gac caa Asp Gln 775	ggt cta Gly Lev	aca gto Thr Val	gtt ggt Val Gly 780	ttg gaa Leu Glu	cct 2352 Pro
tcg tgc ac Ser Cys Th 785					Glu Leu		
cct gat ct Pro Asp Le	cg gca cgc eu Ala Arg 805	ctt gca Leu Ala	gca cto Ala Leu	acc aaa Thr Lys 810	cca tto Pro Phe	gct gag Ala Glu 815	gtc 2448 Val
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aca gaa to Thr Glu Se 83						Arg Ser	
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gat gaa ca Asp Glu G 865					ı Ala Gly		
ttt gaa aa Phe Glu Ly	aa gac cac ys Asp His 885	gct gaa Ala Glu	atg tco Met Sei	ttc gca Phe Ala 890	a ctt ggt a Leu Gly	gaa cga Glu Arg 895	gag 2688 Glu
ctg ttc co Leu Phe Pr	cc aag gtc ro Lys Val 900	aga aaa Arg Lys	gca gaa Ala Glu 905	Gly His	gtg att Val Ile	gct gac Ala Asp 910	ggt 2736 Gly
ttc tcc tc Phe Ser C						Gln Ala	
cac ctt go His Leu A 930	ca gag gtg la Glu Val	gtc tta Val Leu 935	Ser Ile	ttg gag Leu Glu	g caa aac g Gln Asn 940	aac atg Asn Met	gca 2832 Ala
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<212> PRT

<213> Corynebacterium glutamicum

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Asp Ala Gly Ile Phe Arg Arg Val Pro Ala Ala Val Ala Glu Pro Glu 35 40 45

Asn Val Glu Gln Ile Arg Asp Ala Ile Ala Val Ala Val Ala Arg Gly
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Trp Ser Val Val Gly Arg Gly Gly Gly Ser Ser Val Ala Gly Asn Ala 65 70 75 80

Ile Gly Glu Gly Leu Ile Ile Asp Thr Ser Arg Tyr Phe Asn Arg Ile 85 90 95

Leu Asp Ile Asp Pro Val Ala Gln Thr Ala Val Val Glu Pro Gly Val 100 105 110

Val Cys Asp Ala Leu Arg Asp Ala Ala Ala Glu Phe Gly Leu Thr Tyr 115 120 125

Gly Pro Asp Pro Ser Thr His Ser Arg Cys Thr Ile Gly Gly Met Val 130 135 140

Ala Asn Asn Ala Cys Gly Ser His Ser Val Ala Phe Gly Thr Ala Ala 145 150 155 160

Glu Asn Leu Val Asp Val Thr Leu Met Leu Ser Asp Gly Arg Glu Val 165 170 175

Thr Val Thr Lys Asp Gly Cys Asp Asp Ala Glu Ile Asn Gln Lys Leu 180 185 190

Thr Asp Leu Ala Ser Lys Asn Gln Asp Leu Ile Ser Lys Glu Leu Gly
195 200 205

Arg Phe Pro Arg Gln Val Ser Gly Tyr Gly Leu His Tyr Leu Ala His 210 215 220

Asp Met Ala Lys Ala Met Ala Gly Thr Glu Gly Thr Ile Gly Ile Ile 225 230 235 240

Thr Arg Leu Thr Val Lys Leu Val Pro Thr Pro Lys Val Lys Ala Leu 245 250 255

Ala Val Leu Ala Phe Asp Thr Val Phe Asp Ala Ala Arg Ala Ala Ala 260 265 270

Lys Leu Arg Leu Pro Gly Val Ala Thr Ile Glu Gly Met Gly Gly Asp 275 280 285

Leu Leu Ala Ala Leu Arg Ser Lys Gln Gly Gln Ser Glu Ala Gly Gln 290 295 300

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Cys	Glu	Thr	Gly	Ser 325	Asp	Thr	Leu	Gln	Ala 330	Ala	Val	Gln	Ala	Ala 335	Glu
Glu	Val	Ala	Thr 340	Ala	Val	Asp	Thr	Ile 345	Asp	Tyr	Val	Val	Val 350	Ser	Glu
Pro	Ser	Glu 355	Met	Arg	Glu	Leu	Trp 360	Arg	Ile	Arg	Glu	Ser 365	Ser	Ala	Gly
Ile	Val 370	Thr	Arg	Leu	Ala	Asp 375	Gly	Gly	Glu	Ala	Trp 380	Pro	Asn	Trp	Glu
Asp 385	Ser	Ala	Val	Pro	Pro 390	Glu	Asn	Leu	Ala	Asp 395	Tyr	Leu	Arg	Asp	Leu 400
Tyr	Ala	Leu	Met	Asp 405	Lys	Phe	Asp	Tyr	Gln 410	Gly	Ile	Pro	Phe	Gly 415	His
Phe	Gly	Glu	Gly 420	Суѕ	Val	His	Val	Arg 425	Ile	Ser	Phe	Asp	Phe 430	Ser	Thr
Lys	Glu	Gly 435	Leu	Lys	Lys	Phe	Glu 440	Ala	Phe	Met	Asn	Glu 445	Ala	Ser	Thr
Leu	Val 450	Ala	Ser	Tyr	Gly	Gly 455	Ser	Leu	Ser	Gly	Glu 460	His	Gly	Asp	Gly
Arg 465	Ala	Arg	Ser	Ser	Phe 470	Leu	Asp	Arg	Met	Tyr 475	Ser	Ala	Glu	Met	Arg 480
465			Ser Glu		470					475					480
465 Ala	Leu	Phe		Glu 485	470 Phe	Lys	Leu	Ile	Phe 490	475 Asp	Pro	Gln	Arg	Ile 495	480 Phe
465 Ala Asn	Leu Pro	Phe Gly	Glu Val	Glu 485 Leu	470 Phe Val	Lys Trp	Leu Ala	Ile Asp 505	Phe 490 Pro	475 Asp Val	Pro Met	Gln Gln	Arg Gly 510	Ile 495 Leu	480 Phe Arg
465 Ala Asn Met	Leu Pro Asp	Phe Gly Pro 515	Glu Val 500	Glu 485 Leu Gln	470 Phe Val Arg	Lys Trp Ala	Leu Ala Leu 520	Ile Asp 505 Asp	Phe 490 Pro	475 Asp Val Thr	Pro Met Pro	Gln Gln Val 525	Arg Gly 510 His	Ile 495 Leu Lys	480 Phe Arg Phe
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Asn Met Ser Val	Leu Pro Asp Lys 530 Ser	Phe Gly Pro 515 Asp	Glu Val 500 Gly Lys	Glu 485 Leu Gln Gly	470 Phe Val Arg Ser	Lys Trp Ala Met 535 Glu	Leu Ala Leu 520 Ile Ser	Ile Asp 505 Asp Asn	Phe 490 Pro Ile Ala	475 Asp Val Thr Val Met 555	Pro Met Pro Asn 540 Cys	Gln Gln Val 525 Arg	Arg Gly 510 His Cys	Ile 495 Leu Lys Val	Arg Phe Gly Gln 560
Asn Met Ser Val 545	Leu Pro Asp Lys 530 Ser	Phe Gly Pro 515 Asp Ala	Glu Val 500 Gly Lys Cys	Glu 485 Leu Gln Gly Arg	470 Phe Val Arg Ser Ser 550 Val	Lys Trp Ala Met 535 Glu His	Leu Ala Leu 520 Ile Ser	Ile Asp 505 Asp Asn Asp	Phe 490 Pro Ile Ala Ala Arg 570	475 Asp Val Thr Val Met 555 Gly	Pro Met Pro Asn 540 Cys	Gln  Val 525  Arg  Pro	Arg Gly 510 His Cys Ser	Ile 495 Leu Lys Val Phe Leu 575	Arg Phe Gly Gln 560 Leu
Asn Met Ser Val 545 Ile Ser	Leu Pro Asp Lys 530 Ser Thr	Phe Gly Pro 515 Asp Ala Gly Met	Glu Val 500 Gly Lys Cys Asp	Glu 485 Leu Gln Gly Arg Glu 565	470 Phe Val Arg Ser Ser 550 Val	Lys Trp Ala Met 535 Glu His	Leu Ala Leu 520 Ile Ser Ser	Ile Asp 505 Asp Asn Asp Thr	Phe 490 Pro Ile Ala Ala Arg 570 Ala	475 Asp Val Thr Val Met 555 Gly Asp	Pro Met Pro Asn 540 Cys Arg	Gln  Val 525  Arg  Pro Ala	Arg Gly 510 His Cys Ser Arg	Ile 495 Leu Lys Val Phe Leu 575 Ser	Arg Phe Gly Gln 560 Leu Glu

PCT/IB00/00943 WO 01/00844

Leu 625	Asp	Lys	His	Tyr	Ala 630	Gly	Arg	Leu	Arg	Pro 635	Met	Ala	His	Tyr	Val 640
Met	Gly	Trp	Leu	Pro 645	Leu	Leu	Gly	His	Val 650	Ala	His	Lys	Ile	Pro 655	Leu
Leu	Pro	Thr	Leu 660	Ile	Asp	Ala	Thr	Met 665	Gln	Ser	Ala	Leu	Thr 670	Ala	Pro
Val	Val	Arg 675	Lys	Val	Gly	Gly	Leu 680	Ala	Asp	Arg	Pro	Leu 685	Ile	Ser	Phe
Ala	His 690	Arg	Ser	Leu	Arg	Lys 695	Tyr	Lys	Pro	Lys	Lys 700	Asn	Ser	Gly	Glu
Thr 705	Val	Val	Leu	Trp	Pro 710	Asp	Ser	Phe	Asn	Thr 715	Asn	Leu	Asp	Thr	Gly 720
Pro	Ala	His	Ala	Ala 725	Ile	Lys	Thr	Leu	Glu 730	Ala	Leu	Gly	Tyr	Asn 735	Val
Val	Ile	Pro	Asp 740	Gly	Phe	Val	Cys	Cys 745	Gly	Leu	Thr	Trp	His 750	Ser	Thr
Gly	Gln	Leu 755	Ser	Met	Thr	Lys	Lys 760	Val	Leu	Glu	Gln	Thr 765	Ala	Lys	Val
Met	Lys 770	Pro	Tyr	Leu	Asp	Gln 775	Gly	Leu	Thr	Val	Val 780	Gly	Leu	Glu	Pro
Ser 785	Суѕ	Thr	Val	Met	Leu 790	Gln	Asp	Glu	Ala	Thr 795	Glu	Leu	Ser	Asp	Asn 800
Pro	Asp	Leu	Ala	Arg 805	Leu	Ala	Ala	Leu	Thr 810	Lys	Pro	Phe	Ala	Glu 815	Val
Ile	Ala	Pro	Lys 820	Ile	Thr	Glu	Leu	Val 825	Glu	Ser	Gly	Ser	Leu 830	Gln	Leu
Thr	Glu	Ser 835	Thr	Ala	Leu	Thr	Gln 840	Val	His	Cys	His	Glu 845	Arg	Ser	Leu
Gly	Asp 850	Pro	Gln	Gln	Ser	Ala 855	Leu	Val	Leu	Glu	Ala 860	Leu	Gly	Val	Lys
Asp 865	Glu	Gln	Ile	Ala	Thr 870	Gly	Cys	Cys	Gly	Leu 875	Ala	Gly	Asn	Trp	Gly 880
Phe	Glu	Lys	Asp	His 885	Ala	Glu	Met	Ser	Phe 890	Ala	Leu	Gly	Glu	Arg 895	Glu
Leu	Phe	Pro	Lys 900	Val	Arg	Lys	Ala	Glu 905	Gly	His	Val	Ile	Ala 910	Asp	Gly
Phe	Ser	Cys 915	Arg	Thr	Gln	Ile	Glu 920	Gln	Gly	Thr	Gly	Lys 925	Gln	Ala	Thr
His	Leu 930	Ala	Glu	Val	Val	Leu 935	Ser	Ile	Leu	Glu	Gln 940	Asn	Asn	Met	Ala
Gln															

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ctg ccc aac ccc gca gaa cta ctc gaa ctc atg aag ttc aaa aag cca 163 Leu Pro Asn Pro Ala Glu Leu Leu Glu Leu Met Lys Phe Lys Lys Pro 10 15 20	;
gag ctc aac ggc aag aaa cga cgc cta gac tcc gcg ctc acc atc tac 211 Glu Leu Asn Gly Lys Lys Arg Arg Leu Asp Ser Ala Leu Thr Ile Tyr 25 30 35	•
gac ctg cgt aaa att gct aaa cga cgc acc cca gct gcc gcg ttc gac 259 Asp Leu Arg Lys Ile Ala Lys Arg Arg Thr Pro Ala Ala Ala Phe Asp 40 45 50	)
tac acc gac ggc gca gcc gag gcc gaa ctc tca atc aca cgc gca cgt  Tyr Thr Asp Gly Ala Ala Glu Ala Glu Leu Ser Ile Thr Arg Ala Arg  55 60 65	,
gaa gca'ttc gaa aac atc gaa ttc cac cca gac atc ctc aag cct gca 355 Glu Ala Phe Glu Asn Ile Glu Phe His Pro Asp Ile Leu Lys Pro Ala 70 75 80 85	j
gaa cac gta gac acc acc caa atc ctg ggc gga acc tcc tcc atg 403 Glu His Val Asp Thr Thr Thr Gln Ile Leu Gly Gly Thr Ser Ser Met 90 95 100	ţ
cca ttc ggc atc gca cca acc ggc ttc acc cgc ctc atg cag acc gaa 451 Pro Phe Gly Ile Ala Pro Thr Gly Phe Thr Arg Leu Met Gln Thr Glu 105 110 115	•
ggt gaa atc gca ggt gcc gga gct gca ggc gct gca gga att cct ttc 499 Gly Glu Ile Ala Gly Ala Gly Ala Ala Gly Ala Ala Gly Ile Pro Phe 120 125 130	<b>)</b>
acc ctg tcc acc ctg ggc act acc tcc atc gaa gac gtc aag gcc acc  Thr Leu Ser Thr Leu Gly Thr Thr Ser Ile Glu Asp Val Lys Ala Thr  135  140  145	1
aac ccc aac ggc cga aac tgg ttc cag ctc tac gtc atg cgc gac cgc Asn Pro Asn Gly Arg Asn Trp Phe Gln Leu Tyr Val Met Arg Asp Arg 150 155 160 165	j
gaa atc tee tae gge ete gte gaa ege gea gee aaa gea gga tte gae 643	3

Glu	Ile	Ser	Tyr	Gly 170	Leu	Val	Glu	Arg	Ala 175	Ala	Lys	Ala	Gly	Phe 180	Asp	
acc Thr	ctg Leu	atg Met	ttc Phe 185	acc Thr	gtg Val	gat Asp	acc Thr	ccc Pro 190	atc Ile	gcc Ala	ggc Gly	tac Tyr	cgc Arg 195	atc Ile	cgc Arg	691
								ccg Pro								739
gtg Val	ctc Leu 215	aat Asn	gca Ala	atc Ile	cca Pro	cgc Arg 220	cca Pro	tgg Trp	tgg Trp	tgg Trp	atc Ile 225	gac Asp	ttc Phe	ctg Leu	acc Thr	787
acc Thr 230	cca Pro	acc Thr	ctt Leu	gag Glu	ttc Phe 235	gca Ala	tcc Ser	ctt Leu	tcc Ser	tcg Ser 240	acc Thr	ggc Gly	gga Gly	acc Thr	gtg Val 245	835
ggc Gly	gac Asp	ctc Leu	ctc Leu	aac Asn 250	tcc Ser	gcg Ala	atg Met	gat Asp	ccc Pro 255	acc Thr	att Ile	tct Ser	tac Tyr	gaa Glu 260	gac Asp	883
ctc Leu	aag Lys	gtc Val	atc Ile 265	cgt Arg	gaa Glu	atg Met	tgg Trp	cca Pro 270	ggc Gly	aag Lys	ctc Leu	gta Val	gtc Val 275	aag Lys	ggt Gly	931
gtc Val	cag Gln	aac Asn 280	gtt Val	gaa Glu	gac Asp	tcc Ser	gtc Val 285	aaa Lys	ctc Leu	ctc Leu	gac Asp	caa Gln 290	ggc Gly	gtc Val	gac Asp	979
ggc Gly	ctc Leu 295	atc Ile	ctc Leu	tcc Ser	aac Asn	cac His 300	ggt Gly	ggc Gly	cgt Arg	caa Gln	ctc Leu 305	gac Asp	cgc Arg	gca Ala	cca Pro	1027
								gta Val								1075
								atc Ile								1123
gca Ala	gcc Ala	gta Val	gcc Ala 345	atg Met	ggc Gly	gct Ala	gac Asp	ttc Phe 350	acc Thr	ctc Leu	atc Ile	ggt Gly	cgt Arg 355	gcc Ala	tac Tyr	1171
ctc Leu	tac Tyr	gga Gly 360	ctc Leu	atg Met	gcc Ala	gga Gly	ggc Gly 365	cgc Arg	gaa Glu	ggc Gly	gtc Val	gac Asp 370	cgc Arg	acc Thr	atc Ile	1219
Ala								cgc Arg								1267
tcc	Ile 375 tcc	Leu ctc	Arg	Ser gaa	Glu	Ile 380 gag	Thr		Thr	Met gtc	Ala 385 acc	Leu	Leu ctg	Gly	Val aag	1315

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<400> 136

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Ala Leu Thr Ile Tyr Asp Leu Arg Lys Ile Ala Lys Arg Arg Thr Pro
35 40 45

Ala Ala Ala Phe Asp Tyr Thr Asp Gly Ala Ala Glu Ala Glu Leu Ser 50 60

Ile Thr Arg Ala Arg Glu Ala Phe Glu Asn Ile Glu Phe His Pro Asp
65 70 75 80

Ile Leu Lys Pro Ala Glu His Val Asp Thr Thr Thr Gln Ile Leu Gly 85 90 95

Gly Thr Ser Ser Met Pro Phe Gly Ile Ala Pro Thr Gly Phe Thr Arg 100 105 110

Leu Met Gln Thr Glu Gly Glu Ile Ala Gly Ala Gly Ala Ala Gly Ala 115 120 125

Ala Gly Ile Pro Phe Thr Leu Ser Thr Leu Gly Thr Thr Ser Ile Glu 130 135 140

Asp Val Lys Ala Thr Asn Pro Asn Gly Arg Asn Trp Phe Gln Leu Tyr 145 150 155 160

Val Met Arg Asp Arg Glu Ile Ser Tyr Gly Leu Val Glu Arg Ala Ala 165 170 175

Lys Ala Gly Phe Asp Thr Leu Met Phe Thr Val Asp Thr Pro Ile Ala 180 185 190

Gly Tyr Arg Ile Arg Asp Ser Arg Asn Gly Phe Ser Ile Pro Pro Gln 195 200 205

Leu Thr Pro Ser Thr Val Leu Asn Ala Ile Pro Arg Pro Trp Trp 210 215 220

Ile Asp Phe Leu Thr Thr Pro Thr Leu Glu Phe Ala Ser Leu Ser Ser 225 230 235 240

Thr Gly Gly Thr Val Gly Asp Leu Leu Asn Ser Ala Met Asp Pro Thr 245 250 255

Ile Ser Tyr Glu Asp Leu Lys Val Ile Arg Glu Met Trp Pro Gly Lys 260 265 270

Leu Val Val Lys Gly Val Gln Asn Val Glu Asp Ser Val Lys Leu Leu Asp Gln Gly Val Asp Gly Leu Ile Leu Ser Asn His Gly Gly Arg Gln Leu Asp Arg Ala Pro Val Pro Phe His Leu Leu Pro Gln Val Arg Lys 315 Glu Val Gly Ser Glu Pro Thr Ile Met Ile Asp Thr Gly Ile Met Asn Gly Ala Asp Ile Val Ala Ala Val Ala Met Gly Ala Asp Phe Thr Leu Ile Gly Arg Ala Tyr Leu Tyr Gly Leu Met Ala Gly Gly Arg Glu Gly Val Asp Arg Thr Ile Ala Ile Leu Arg Ser Glu Ile Thr Arg Thr Met 375 Ala Leu Leu Gly Val Ser Ser Leu Glu Glu Leu Glu Pro Arg His Val 390 395 Thr Gln Leu Ala Lys Met Val Pro Val Ser Asp Ala Thr Arg Ser Ala 405 410 Ala Ala Glu Ile 420 <210> 137 <211> 1836 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(1813) <223> RXN01952 <400> 137 ccatcaaaaa atqaacqacc qcqqactaqc tcggatcaag gcgacatccc ctcagcatca 60 tqacqcqctt qtqatqcaac tqaatataqg aagcttaqag atg acg caa cca gga Met Thr Gln Pro Gly cag acc acc acg act tcg cac gaa gcg atc gat gcg ttc aag aga atc 163 Gln Thr Thr Thr Thr Ser His Glu Ala Ile Asp Ala Phe Lys Arg Ile gtc ggc gac gaa cat gta ctg acc tct gag cgt gcc acg atg cca ttc 211 Val Gly Asp Glu His Val Leu Thr Ser Glu Arg Ala Thr Met Pro Phe age aaa ggc tat cga tte ggc gga gga cca gte tte gee gtg gtg ege 259 Ser Lys Gly Tyr Arg Phe Gly Gly Gly Pro Val Phe Ala Val Val Arg 45 40

ccc Pro	ggc Gly 55	acg Thr	ctg Leu	gtc Val	gag Glu	atg Met 60	tgg Trp	cgg Arg	gcg Ala	ctg Leu	cag Gln 65	gta Val	tcc Ser	gtc Val	gac Asp	307
	aac Asn															355
gga Gly	tcc Ser	ggc Gly	ccc Pro	ggc Gly 90	ttc Phe	caa Gln	gac Asp	tac Tyr	gat Asp 95	cgc Arg	ccc Pro	att Ile	gtg Val	atc Ile 100	atc Ile	403
	act Thr															451
gcg Ala	atc Ile	tcg Ser 120	ctc Leu	gcg Ala	ggc Gly	acc Thr	ccg Pro 125	ctg Leu	aca Thr	cac His	ctg Leu	acc Thr 130	gac Asp	gcg Ala	ctc Leu	499
gcc Ala	aag Lys 135	cac His	cag Gln	cgc Arg	gag Glu	ccg Pro 140	cac His	tcg Ser	gtg Val	atc Ile	ggg Gly 145	tcg Ser	aca Thr	tca Ser	atc Ile	547
	gcc Ala															595
	.cgc Arg															643
	gac Asp															691
	gac Asp															739
	ccc Pro 215			Val		Pro	Ala	Pro	Glu		Ser	Asn				787
	gcc Ala															835
	aac Asn															883
	gtg Val															931
	gtg Val															979
cgt	cgg	ttg	ttc	ctc	gaa	gcc	gac	atg	ccg	ctg	cct	atc	tct	ggt	gag	1027

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acc Thr	ttc Phe	gtc Val	ttc Phe	ctg Leu 330	aag Lys	ttc Phe	atg Met	agt Ser	cca Pro 335	gcg Ala	ctg Leu	cag Gln	acg Thr	cgc Arg 340	atg Met	1123
ttc Phe	tcg Ser	ttc Phe	aag Lys 345	acg Thr	tgg Trp	gcc Ala	aac Asn	ggc Gly 350	ttg Leu	ttc Phe	tcg Ser	aag Lys	att Ile 355	ccc Pro	ggc Gly	1171
	ggt Gly															1219
ctg Leu	ccc Pro 375	aac Asn	cag Gln	ctg Leu	ccc Pro	aag Lys 380	cgc Arg	atg Met	atg Met	gag Glu	tac Tyr 385	cgc Arg	aac Asn	cgt Arg	ttc Phe	1267
	cat His															1315
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	ttc Phe		-	_		-	•	-	_	-		_				1411
	ggc Gly															1459
	atc Ile 455															1507
tgg Trp 470	aac Asn	tgg Trp	ctc Leu	gag Glu	gtg Val 475	ctg Leu	ccg Pro	gag Glu	gag Glu	atc Ile 480	gac Asp	gac Asp	cag Gln	ctt Leu	gag Glu 485	1555
	aag Lys															1603
	gtc Val															1651
cag Gln	cac His	ctg Leu 520	ctg Leu	gag Glu	gag Glu	cgc Arg	ggc Gly 525	gcg Ala	aag Lys	ctg Leu	ccc Pro	gcc Ala 530	gag Glu	cac His	aac Asn	1699
	ggt Gly															1747

535 540 545

gag ctc gat ccg acg aat acg ttc aac gcc ggt atc ggc ggc acg tcg 1795 Glu Leu Asp Pro Thr Asn Thr Phe Asn Ala Gly Ile Gly Gly Thr Ser 550 565

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Ala Thr Met Pro Phe Ser Lys Gly Tyr Arg Phe Gly Gly Gly Pro Val

Phe Ala Val Val Arg Pro Gly Thr Leu Val Glu Met Trp Arg Ala Leu 50 55 60

Gln Val Ser Val Asp Asn Asn Leu Ile Val Ile Pro Gln Ala Ser Asn 65 70 75 80

Thr Gly Leu Thr Gly Gly Ser Gly Pro Gly Phe Gln Asp Tyr Asp Arg 85 90 95

Pro Ile Val Ile Ile Ser Thr His Arg Ile Asp Glu Val His Leu Ile 100 105 110

Asn Asp Ala Arg Glu Ala Ile Ser Leu Ala Gly Thr Pro Leu Thr His 115 120 125

Leu Thr Asp Ala Leu Ala Lys His Gln Arg Glu Pro His Ser Val Ile 130 135 140

Gly Ser Thr Ser Ile Gly Ala Ser Val Ile Gly Gly Ile Ala Asn Asn 145 150 155 160

Ser Gly Gly Ser Gln Ile Arg Lys Gly Pro Ala Phe Thr Arg Glu Ala 165 170 175

Ile Phe Ala Arg Val Asn Asp Asp Gly Lys Val Glu Leu Val Asn His 180 185 190

Leu Gly Ile Ser Leu Gly Asp Asp Pro Glu Val Ala Leu Asp Arg Leu
195 200 205

Gln Arg Gly Glu Trp Ser Pro Glu Asp Val Thr Pro Ala Pro Glu Asp 210 215 220

Ser Asn Glu Thr Glu Tyr Ala Glu His Leu Arg Lys Ile Val Pro Ser 225 230 235 240

Pro	Ala	Arg	Tyr	Asn 245	Ala	Asn	Pro	Glu	Tyr 250	Leu	Phe	Glu	Ala	Ser 255	Gly
Ser	Ala	Gly	Lys 260	Leu	Met	Val	Phe	Ala 265	Val	Arg	Thr	Arg	Thr 270	Phe	Pro
Arg	Glu	Val 275	His	Pro	Thr	Val	Phe 280	Tyr	Ile	Gly	Thr	Asn 285	Asn	Thr	His
Glu	Leu 290	Glu	Glu	Ile	Arg	Arg 295	Leu	Phe	Leu	Glu	Ala 300	Asp	Met	Pro	Leu
Pro 305	Ile	Ser	Gly	Glu	Tyr 310	Met	Gly	Arg	Ser	Ala 315	Phe	Asp	Leu	Ala	Glu 320
Lys	Tyr	Gly	Lys	Asp 325	Thr	Phe	Val	Phe	Leu 330	Lys	Phe '	Met	Ser	Pro 335	Ala
Leu	Gln	Thr	Arg 340	Met	Phe	Ser	Phe	Lys 345	Thr	Trp	Ala	Asn	Gly 350	Leu	Phe
Ser	Lys	Ile 355	Pro	Gly	Ile	Gly	Pro 360	Thr	Phe	Ala	Asp	Thr 365	Val	Ser	Gln
Ala	Met 370	Phe	Ser	Val	Leu	Pro 375	Asn	Gln	Leu	Pro	Lys 380	Arg	Met	Met	Glu
Tyr 385	Arg	Asn	Arg	Phe	Glu 390	His	His	Leu	Leu	Leu 395	Thr	Val	Ser	Glu	Ser 400
Gln	Lys	Ala	Ala	Ser 405	Glu	Lys	Met	Leu	Lys 410	Glu	Phe	Phe	Ala	Glu 415	Pro
Glu	His	Thr	Gly 420	Glu	Phe	Phe	Ile	Cys 425	Thr	Ser	Asp	Glu	Glu 430	Lys	Ser
Ala	Ser	Leu 435	Asn	Arg	Phe	Gly	Ala 440	Ala	Ser	Ala	Ala	Thr 445	Arg	Tyr	Ala
Ala	Leu 450	Lys	Arg	Arg	His	Ile 455	Ala	Gly	Leu	Ile	Pro 460	Ile	Asp	Val	Ala
Leu 465	Arg	Arg	Asp	Asp	Trp 470	Asn	Trp	Leu	Glu	Val 475	Leu	Pro	Glu	Glu	Ile 480
Asp	Asp	Gln	Leu	Glu 485	Val	Lys	Ala	Tyr	Tyr 490	Gly	His	Phe	Phe	Cys 495	His
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Leu	His	Asp 515	Arg	Ile	Gln	His	Leu 520	Leu	Glu	Glu	Arg	Gly 525	Ala	Lys	Leu
Pro	Ala 530	Glu	His	Asn	Tyr	Gly 535	Arg	Met	Tyr	Lys	Leu 540	Pro	Glu	Ser	Met
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<223> FRXA01952

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cac aac tac ggt cgc atg tac aag ctg ccg gag tcc atg gaa gag cac  $\phantom{0}$  144 His Asn Tyr Gly Arg Met Tyr Lys Leu Pro Glu Ser Met Glu Glu His  $\phantom{0}$  35  $\phantom{0}$  40  $\phantom{0}$  45

ttc aag gag ctc gat ccg acg aat acg ttc aac gcc ggt atc ggc ggc 192
Phe Lys Glu Leu Asp Pro Thr Asn Thr Phe Asn Ala Gly Ile Gly Gly
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Thr Ser Pro His Lys Asp Trp Ala
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<400> 140

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His Asn Tyr Gly Arg Met Tyr Lys Leu Pro Glu Ser Met Glu Glu His 35 40 45

Phe Lys Glu Leu Asp Pro Thr Asn Thr Phe Asn Ala Gly Ile Gly Gly 50 55 60

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190

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tct Ser	ccc Pro 215	gag Glu	gat Asp	gtc Val	acc Thr	cca Pro 220	gct Ala	ccc Pro	gaa Glu	gac Asp	tcg Ser 225	aac Asn	gag Glu	acc Thr	gag Glu	787
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acc Thr	gtg Val	ttt Phe 280	tac Tyr	atc Ile	ggc Gly	acg Thr	aac Asn 285	aac Asn	acg Thr	cac His	gag Glu	ctc Leu 290	gaa Glu	gag Glu	atc Ile	979
cgt Arg	cgg Arg 295	ttg Leu	ttc Phe	ctc Leu	gaa Glu	gcc Ala 300	gac Asp	atg Met	ccg Pro	ctg Leu	cct Pro 305	atc Ile	tct Ser	ggt Gly	gag Glu	1027
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					tgg Trp											1171
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					ctc Leu 395											1315
					gag Glu											1363
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gtc /al	aag Lys	gcg Ala	tat Tyr	tac Tyr 490	Gly	cac His	ttc Phe	ttc Phe	tgc Cys 495	cat His	gtg Val	atg Met	cac His	cag Gln 500	gac Asp	1603
at Tyr	gtc Val	gcc Ala	aag Lys 505	cag Gln	ggc Gly	gtg Val	gat Asp	ctc Leu 510	gag Glu	gcg Ala	ctg Leu	cac His	gac Asp 515	cgc Arg	atc Ile	1651
cag Gln	cac His	ctg Leu 520	ctg Leu	gag Glu	gag Glu	cgc Arg	ggc Gly 525	gcg Ala	aag Lys	ctg Leu	ccc Pro	gcc Ala 530	gag Glu	cac His	aac Asn	1699
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Ala	Thr	Met 35	Pro	Phe	Ser	Lys	Gly 40	Tyr	Arg	Phe	Gly	Gly 45	Gly	Pro	Val	
Phe	Ala 50	Val	Val	Arg	Pro	Gly 55	Thr	Leu	Val	Glu	Met 60	Trp	Arg	Ala	Leu	
Gln 65	Val	Ser	Val	Asp	Asn 70	Asn	Leu	Ile	Val	Ile 75	Pro	Gln	Ala	Ser	Asn 80	
	ctc Phe cassis cgg property can be considered to the cassis cgg property can be considered to the cassis considered to th	tc ggc Phe Gly cac atc lis Ile 455 cgg aac Trp Asn 170 gtc aag Val Lys cat gtc Tyr Val cag cac Gln His <210> 14 <211> 53 <212> Pi <213> Co <400> 14 Ala Phe Ala Thr Phe Ala 50 Gln Val	tc ggc gcg Phe Gly Ala 440 cac atc gca dis Ile Ala 455 cgg aac tgg Trp Asn Trp Tro gtc aag gcg Val Lys Ala cat gtc gcc Tyr Val Ala cag cac ctg Gln His Leu 520 c210> 142 c211> 533 c212> PRT c213> Coryn c400> 142 Met Thr Gln 1 Ala Phe Lys Ala Thr Met 35 Phe Ala Val 50 Gln Val Ser	Che Gly Ala Ala  440  Cac atc gca ggg  Iis Ile Ala Gly  455  Cgg aac tgg ctc  Crp Asn Trp Leu  170  Gtc aag gcg tat  7al Lys Ala Tyr  Cat gtc gcc aag  Tyr Val Ala Lys  505  Cag cac ctg ctg  Gln His Leu Leu  520  C210> 142  C211> 533  C212> PRT  C213> Corynebac  C400> 142  Met Thr Gln Pro  1  Ala Phe Lys Arg  20  Ala Thr Met Pro  35  Phe Ala Val Val  50  Gln Val Ser Val	the ggc gcg gcc agt Phe Gly Ala Ala Ser 440  cac atc gca ggg ctc dis Ile Ala Gly Leu 455  agg aac tgg ctc gag Asn Trp Leu Glu Afro  gtc aag gcg tat tac Val Lys Ala Tyr Tyr 490  cat gtc gcc aag cag Tyr Val Ala Lys Gln 505  cag cac ctg ctg gag Gln His Leu Leu Glu 520  c210> 142 c211> 533 c212> PRT c213> Corynebacteriu c400> 142 Met Thr Gln Pro Gly 1 Shala Phe Lys Arg Ile 20  Ala Thr Met Pro Phe 35  Phe Ala Val Val Arg 50  Gln Val Ser Val Asp	che Gly Ala Ala Ser Ala 440  cac atc gca ggg ctc atc lis Ile Ala Gly Leu Ile 455  cgg aac tgg ctc gag gtg Trp Asn Trp Leu Glu Val 475  gtc aag gcg tat tac ggg Yal Lys Ala Tyr Tyr Gly 490  cat gtc gcc aag cag ggc Tyr Val Ala Lys Gln Gly 505  cag cac ctg ctg gag gag Gln His Leu Leu Glu Glu 520  c210> 142 c211> 533 c212> PRT c213> Corynebacterium g c400> 142 Met Thr Gln Pro Gly Gln 1 5  Ala Phe Lys Arg Ile Val 20  Ala Thr Met Pro Phe Ser 35  Phe Ala Val Val Arg Pro 50  Gln Val Ser Val Asp Asn	the ggc gcg gcc agt gcc gcc  The Gly Ala Ala Ser Ala Ala  A440  The Gly Ala Cyc atc gcc  The Ala Gly Leu Ile Pro  A55  The Ala Gly Leu Ile Pro  A55  The Ala Cyc gcc gag gtg ctg  The Gly Ala Tyr Tyr Gly His  A90  The Gly Ala Tyr Tyr Gly His  A90  The Ala Leu Leu Glu Glu Arg  The Gly Cyc agg gag cyc  The Gly Ala Leu Glu Glu Arg  The Cyc ala Cyc gag gag cyc  The Ala Cyc ala Cyc gag gag cyc  The Gly Cyc ala Cy	the ggc gcg gcc agt gcc gcc act he Gly Ala Ala Ser Ala Ala Thr 440  aca atc gca ggg ctc atc ccc atc lis Ile Ala Gly Leu Ile Pro Ile 455  agg aac tgg ctc gag gtg ctg ccg frp Asn Trp Leu Glu Val Leu Pro 475  gtc aag gcg tat tac ggg cac ttc Tyr Gly His Phe 490  at gtc gcc aag cag ggc gtg gat fyr Val Ala Lys Gln Gly Val Asp 505  agg cac ctg ctg gag gag cgc ggc gat Gly Val Asp 505  agg cac ctg ctg gag gag cgc ggc gar Gln His Leu Leu Glu Glu Arg Gly 525  agg cac ctg ctg gag gag cgc ggc gag cac ttc Gly S20  ala Phe Lys Arg Ile Val Gly Asp 20  Ala Thr Met Pro Phe Ser Lys Gly 35  Phe Ala Val Val Arg Pro Gly Thr 50  Gln Val Ser Val Asp Asn Asn Leu	the ggc gcg gcc agt gcc act cgc Ala Ala Thr Arg 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tcgc Val Lys Ala Tyr Tyr Gly His Phe Phe Cys 490  act gtc gcc aag cag ggc gtg gat ctc gag Gry Val Ala Lys Gln Gly Val Asp Leu Glu 510  acag cac ctg ctg gag gag cgc ggc gcg aag Gln His Leu Leu Glu Glu Arg Gly Ala Lys 520  act Cat Cat ctg gag gag cgc gtg gcc act ctc tact gag Gln His Leu Leu Glu Glu Arg Gly Ala Lys 520  act gtc Scall > 142  act Thr Gln Pro Gly Gln Thr Thr Thr Thr 1 1 1 10  ala Phe Lys Arg Ile Val Gly Asp Glu His 20  ala Thr Met Pro Phe Ser Lys Gly Tyr Arg 35  act Cat Val Asp Asn Asn Leu Ile Val	the ggc gcg gcc agt gcc act cgc tac gcc he Gly Ala Ala Ser Ala Ala Thr Arg Tyr Ala 440  The Gly Ala Ala Ser Ala Ala Thr Arg Tyr Ala 445  The Gly Ala Ala Ser Ala Ala Thr Arg Tyr Ala 445  The Ala Gly Leu Ile Pro Ile Asp Val Ala 455  The Ala Gly Leu Ile Pro Ile Asp Val Ala 460  The Ash Trp Leu Glu Val Leu Pro Glu Glu Ile 475  The Ash Trp Leu Glu Val Leu Pro Glu Glu Ile 475  The Ash Trp Leu Glu Val Leu Pro Glu Glu Ile 480  The Ash Trp Leu Glu Val His Phe Phe Cys His 490  The Ala Lys Ala Tyr Tyr Gly His Phe Phe Cys His 490  The Ala Val Val Glu Arg Gly Ala Lys Leu 525  The Ala Val Val Arg Pro Gly Thr Leu Val Glu File Ala Val Val Arg Pro Gly Thr Leu Val Glu Glu Arg Cly Tyr Arg Phe Ala Val Ser Val Asp Ash Ash Leu Ile Val Ile	the ggc gcg gcc agt gcc gcc act cgc tac gcc gcg the Gly Ala Ala Ser Ala Ala Thr Arg Tyr Ala Ala Ala 445  aca atc gca ggg ctc atc ccc atc gat gtg gcc ctg lie Ile Ala Gly Leu Ile Pro Ile Asp Val Ala Leu 465  agg aac tgg ctc gag gtg ctg ccg gag gag atc gac trop Asn Trp Leu Glu Val Leu Pro Glu Glu Ile Asp 480  act aag gcg tat tac ggg cac ttc ttc tgc cat gtg val Lys Ala Tyr Tyr Gly His Phe Phe Cys His Val 490  act gtc gcc aag cag ggc gtg gat ctc gag gcg ctg yv Val Ala Leu 505  act gtc gcc aag cag ggc gtg gat ctc gag gcg ctg val Ala Leu 505  act gtc gcc aag cag gag cgc ggc gcg aag ctg ccc glu Ala Leu 500  act gtc gcc aag cag gag cgc ggc gcg aag ctg ccc gal His Leu Leu Glu Glu Arg Gly Ala Lys Leu Pro 525  act cac ctg ctg gag gag cgc ggc gcg aag ctg ccc 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the Gly Ala Ala Ser Ala Ala Thr Arg Tyr Ala Ala Leu 440  aca atc gca ggg ctc atc ccc atc gat gtg gcc ctg cgt lis Ile Ala Gly Leu Ile Pro Ile Asp Val Ala Leu 455  acg aac tgg ctc gag gtg ctg ccg gag gag atc gac frp Asn Trp Leu Glu Val Leu Pro Glu Glu Ile Asp Asp 480  act aag gcg tat tac ggg cac ttc ttc tgc cat gtg atg fro Ala Lys Ala Tyr Tyr Gly His Phe Phe Cys His Val Met 490  act gcc aag gcg gtg gat ctc gag gcg gtg gat ctc gag gcg ctg cac fry Val Ala Lys Gln Gly Val Asp Leu Glu Ala Leu His 505  acg cac ctg ctg gag gag cgc ggc gcg aag ctg ccc gcc Sin His Leu Leu Glu Glu Arg Gly Ala Lys Leu Pro Ala 520  act corynebacterium glutamicum  act cory	the ggc gcg gcc agt gcc gcc act cgc tac gcc gcg ttg aag he Gly Ala Ala Ser Ala Ala Thr Arg Tyr Ala Ala Leu Lys 440  act atc gca ggg ctc atc ccc atc gat gtg gcc ctg cgc gcg lis Ile Ala Gly Leu Ile Pro Ile Asp Val Ala Leu Arg Arg 465  acg aac tgg ctc gag gtg ctg ccg gag gag atc gac gac cag grp Asn Trp Leu Glu Val Leu Pro Glu Glu Ile Asp Asp Gln 475  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Thr Gly Leu Thr Gly Gly Ser Gly Pro Gly Phe Gln Asp Tyr Asp Arg 85 90 95

Pro Ile Val Ile Ile Ser Thr His Arg Ile Asp Glu Val His Leu Ile 100 105 110

Asn Asp Ala Arg Glu Ala Ile Ser Leu Ala Gly Thr Pro Leu Thr His 115 120 125

Leu Thr Asp Ala Leu Ala Lys His Gln Arg Glu Pro His Ser Val Ile 130 140

Gly Ser Thr Ser Ile Gly Ala Ser Val Ile Gly Gly Ile Ala Asn Asn 145 150 155 160

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Gln	Arg 210	Gly	Glu	Trp	Ser	Pro 215	Glu	Asp	Val	Thr	Pro 220	Ala	Pro	Glu	Asp
Ser 225	Asn	Glu	Thr	Glu	Tyr 230	Ala	Glu	His	Leu	Arg 235	Lys	Ile	Val	Pro	Ser 240
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Glu	Leu 290	Glu	Glu	Ile	Arg	Arg 295	Leu	Phe	Leu	Glu	Ala 300	Asp	Met	Pro	Leu
Pro 305	Ile	Ser	Gly	Glu	Tyr 310	Met	Gly	Arg	Ser	Ala 315	Phe	Asp	Leu	Ala	Glu 320
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Tyr 385	Arg	Asn	Arg	Phe	Glu 390	His	His	Leu	Leu	Leu 395	Thr	Val	Ser	Glu	Ser 400
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gat gca acg gaa Asp Ala Thr Glu	cct ata tcc Pro Ile Ser 170	aac ctg cac Asn Leu His 175	caa gta ctt gcc Gln Val Leu Ala	gac gcc 643 Asp Ala 180								
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Asp Gly Val Arg Trp Val Gln Phe Pro Asn Ala Gly Leu Asn Ala Tyr Phe Thr Ala Gly Gln Ile Asp Asp Lys Arg Arg Trp Ser Asn Ala Ser Gly Val Tyr Gly Gln Gln Val Ala Glu Ala Ala Met Ala Leu Leu Gly Leu Ile His Met His Pro Thr Met Val Arg Ala Asp Ser Trp Ala Pro Ser Thr Gln Ile Asp Gln Gln Thr Arg Trp Leu Asp Gly Ala Thr 120 Val Ala Ile Val Gly Ala Gly Gly Ile Gly Lys His Leu Ala Ala Met Leu Lys Pro Phe Gly Ala Lys Ser Leu Ala Val Ser Arg Thr Gly Thr 155 150 Pro Thr Gln Asp Phe Asp Ala Thr Glu Pro Ile Ser Asn Leu His Gln 170 Val Leu Ala Asp Ala Asp His Val Val Leu Cys Val Pro Leu Thr Ala 180 Asp Thr Tyr His Leu Ile Gly Lys Ala Glu Leu Lys Ala Met Gln Ser 200 Thr Ala Ile Leu Ile Asn Val Ala Arg Gly Glu Val Val Asp Thr Glu Ala Leu Val Asp Ala Leu Asp Ala Gln Glu Ile Ser Gly Ala Gly Leu Asp Val Thr Asp Pro Glu Pro Leu Pro Asp Asp His Pro Leu Trp Gly Arg Ser Asn Val Ile Ile Thr Pro His Val Ala Asn Thr Leu Thr Ser 265 Met Asp Arg Met Leu Ala Pro Val Val Ala Glu Asn Tyr Arg Arg Phe 280 275

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<sup>&</sup>lt;213> Corynebacterium glutamicum

<sup>&</sup>lt;220>

<sup>&</sup>lt;221> CDS

<sup>&</sup>lt;222> (62)..(664)

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<211> 201

<212> PRT

<213> Corynebacterium glutamicum

<400> 146

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Glu Gln Val Asn Ala Leu Gly Leu Ser Ala Val Arg Gly Leu Phe Ser 35 40 45

Gly Ile Ile Glu Glu Ser Val Thr Phe Val Asn Ala Pro Arg Ile Ala 50 55 60

Glu Glu Arg Gly Leu Asp Ile Ser Val Lys Thr Asn Ser Glu Ser Val 65 70 75 80

Thr His Arg Ser Val Leu Gln Val Lys Val Ile Thr Gly Ser Gly Ala 85 90 95

Ser Ala Thr Val Val Gly Ala Leu Thr Gly Leu Glu Arg Val Glu Lys 100 105 110

Ile Thr Arg Ile Asn Gly Arg Gly Leu Asp Leu Arg Ala Glu Gly Leu 115 120 125

Asn Leu Phe Leu Gln Tyr Thr Asp Ala Pro Gly Ala Leu Gly Thr Val 130 135 140

Gly Thr Lys Leu Gly Ala Ala Gly Ile Asn Ile Glu Ala Ala Ala Leu 145 150 155 160

Thr Gln Ala Glu Lys Gly Asp Gly Ala Val Leu Ile Leu Arg Val Glu 165 170 175

Ser Ala Val Ser Glu Glu Leu Glu Ala Glu Ile Asn Ala Glu Leu Gly 180 185 190

Ala Thr Ser Phe Gln Val Asp Leu Asp 195 200

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<223> FRXA01130

<400> 147

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aat ggc cgt ggc ctg gat ctg cgc gca gag ggt ctg aac ctc ttc ctg 96

Asn Gly Arg Gly Leu Asp Leu Arg Ala Glu Gly Leu Asn Leu Phe Leu cag tac act gac gct cct ggt gca ctg ggt acc gtt ggt acc aag ctg 144 Gln Tyr Thr Asp Ala Pro Gly Ala Leu Gly Thr Val Gly Thr Lys Leu ggt gct gct ggc atc aac atc gag gct gct gcg ttg act cag gct gag 192 Gly Ala Ala Gly Ile Asn Ile Glu Ala Ala Ala Leu Thr Gln Ala Glu 55 240 aag ggt gac ggc gct gtc ctg atc ctg cgt gtt gag tcc gct gtc tct Lys Gly Asp Gly Ala Val Leu Ile Leu Arg Val Glu Ser Ala Val Ser gaa gag ctg gaa gct gaa atc aac gct gag ttg ggt gct act tcc ttc 288 Glu Glu Leu Glu Ala Glu Ile Asn Ala Glu Leu Gly Ala Thr Ser Phe 90 326 cag gtt gat ctt gac taattagaga tccatttgct tga Gln Val Asp Leu Asp 100 <210> 148 <211> 101 <212> PRT <213> Corynebacterium glutamicum <400> 148 Val Gly Ala Leu Thr Gly Leu Glu Arg Val Glu Lys Ile Thr Arg Ile Asn Gly Arg Gly Leu Asp Leu Arg Ala Glu Gly Leu Asn Leu Phe Leu Gln Tyr Thr Asp Ala Pro Gly Ala Leu Gly Thr Val Gly Thr Lys Leu Gly Ala Ala Gly Ile Asn Ile Glu Ala Ala Ala Leu Thr Gln Ala Glu Lys Gly Asp Gly Ala Val Leu Ile Leu Arg Val Glu Ser Ala Val Ser Glu Glu Leu Glu Ala Glu Ile Asn Ala Glu Leu Gly Ala Thr Ser Phe Gln Val Asp Leu Asp 100 <210> 149 <211> 604 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(604) <223> RXN03112

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gctgcgtgag ggcgagtgga agcggtcttc tttcaacggt gtg gaa att ttc gga Val Glu Ile Phe Gly 1 5													
aaa act gtc ggt Lys Thr Val Gly	atc gtc ggt Ile Val Gly 10	ttt ggc cac att Phe Gly His Ile 15	ggt cag ttg ttt gct Gly Gln Leu Phe Ala 20	163									
			gct tac gat cct tac Ala Tyr Asp Pro Tyr 35	211									
	Arg Ala Ala		gag ttg gtt gag ttg Glu Leu Val Glu Leu 50	259									
			att cac ctt cct aag Ile His Leu Pro Lys 65	307									
			ctc ctt gct aag tcc Leu Leu Ala Lys Ser 85	355									
			ggt ggc ctt gtt gat . Gly Gly Leu Val Asp 100	403									
			cac att cgt ggc gct His Ile Arg Gly Ala 115	451									
	Tyr Ser Thr		gat tot cot ttg ttc Asp Ser Pro Leu Phe 130	499									
			ggt gct tct act gaa Gly Ala Ser Thr Glu 145	547									
			gat tot gtg otc aag Asp Ser Val Leu Lys 165	595									
gcg ctg gct Ala Leu Ala				604									
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<400> 150 Val Glu Ile Phe 1	Gly Lys Thr	Val Gly Ile Val	Gly Phe Gly His Ile 15										

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	_			_	gat Asp	_	_	-		_	_	-			-	307
					gcc Ala 75											355
gct Ala	gcc Ala	act Thr	gaa Glu	gct Ala 90	ggc Gly	gtc Val	atg Met	gtt Val	gct Ala 95	aac Asn	gca Ala	ccg Pro	acc Thr	tct Ser 100	aat Asn	403
					gag Glu											451
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-	ggt Gly															649

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<211> 183

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<213> Corynebacterium glutamicum

<400> 152

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Gln Ser Thr Val Asp Ala Leu Gly Asp Ala Val Glu Val Arg Trp Val 20 25 30

Asp Gly Pro Asn Arg Pro Glu Leu Leu Asp Ala Val Lys Glu Ala Asp
35 40 45

Ala Leu Leu Val Arg Ser Ala Thr Thr Val Asp Ala Glu Val Ile Ala 50 55 60

Ala Ala Pro Asn Leu Lys Ile Val Gly Arg Ala Gly Val Gly Leu Asp
65 70 75 80

Asn Val Asp Ile Pro Ala Ala Thr Glu Ala Gly Val Met Val Ala Asn 85 90 95

Ala Pro Thr Ser Asn Ile His Ser Ala Cys Glu His Ala Ile Ser Leu Leu Leu Ser Thr Ala Arg Gln Ile Leu Leu Leu Met Arg Arg Cys Val 120 Arg Ala Ser Gly Ser Gly Leu Leu Ser Thr Val Trp Lys Phe Ser Glu Lys Leu Ser Val Ser Ser Val Leu Ala Thr Leu Val Ser Cys Leu Leu 155 Ser Val Leu Leu Arg Leu Arg Pro Pro Leu Leu Thr Ile Leu Thr 170 Leu Thr Leu Leu Arg Ala Gly 180 <210> 153 <211> 1011 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(988) <223> RXN00871 <400> 153 qqqaaaaggc gatcaccagc cgttggctcg acccagcaac ccacggtggc attaacctcg 60 gtttcccaca gaacgattaa ttgaaggaga gcacaggact atg cgt tgg ttc cat Met Arg Trp Phe His aag aag ggc gaa ctg gcc cga gat ggt tgg caa agc gtt gtc gat gcc Lys Lys Gly Glu Leu Ala Arg Asp Gly Trp Gln Ser Val Val Asp Ala 20 acc acc cca gqt tqq qaa tat acc ggc atc cgc att gcc gaa ctg ggc 211 Thr Thr Pro Gly Trp Glu Tyr Thr Gly Ile Arg Ile Ala Glu Leu Gly 25 30 259 agt ggt gaa tcg ctt gaa ctg aat gac act ggt gtg gaa cgc atc ttc Ser Gly Glu Ser Leu Glu Leu Asn Asp Thr Gly Val Glu Arg Ile Phe 45 40 att cca ctt cag ggc agc ttc gat gtt gcc cac cat ggt cag gtg acc 307 Ile Pro Leu Gln Gly Ser Phe Asp Val Ala His His Gly Gln Val Thr 55 355 cat ctt cac gga aga aag tca gtc ttt gat gga cca acc gat gtg ctc His Leu His Gly Arg Lys Ser Val Phe Asp Gly Pro Thr Asp Val Leu 70 tac etc ecc act gga caa aca gea acg etc agt ggt cag gga ega gte Tyr Leu Pro Thr Gly Gln Thr Ala Thr Leu Ser Gly Gln Gly Arg Val 90 100 gcc gtg gcg gaa gct ccc act cag gaa ccc aag gag tgg aag tac atc

Ala	Val	Ala	Glu 105	Ala	Pro	Thr	Gln	Glu 110	Pro	Lys	Glu	Trp	Lys 115	Tyr	Ile	
gct Ala	cca Pro	gca Ala 120	gaa Glu	act Thr	cct Pro	gtg Val	gag Glu 125	ttg Leu	cgt Arg	gga Gly	gct Ala	ggc Gly 130	cgc Arg	tcg Ser	agc Ser	499
	caa Gln 135															547
	atc Ile															595
	cca Pro															643
	gaa Glu															691
gcc Ala	gaa Glu	gca Ala 200	gca Ala	gaa Glu	gga Gly	gct Ala	ttc Phe 205	gga Gly	atg Met	ttt Phe	tcc Ser	acc Thr 210	tac Tyr	tcc Ser	tca Ser	739
	gcg Ala 215															787
	cta Leu															835
	gac Asp															883
	tgg Trp															931
	acc Thr															979
-	gag Glu 295		taaa	aatti	tca t	ggct	gaaa	ac ga	aa							1011
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	0> 1: Arg		Phe	His 5	Lys	Lys	Gly	Glu	Leu 10	Ala	Arg	Asp	Gly	Trp 15	Gln	

Ser Val Val Asp Ala Thr Thr Pro Gly Trp Glu Tyr Thr Gly Ile Arg
20 25 30

Ile Ala Glu Leu Gly Ser Gly Glu Ser Leu Glu Leu Asn Asp Thr Gly 35 40 45

Val Glu Arg Ile Phe Ile Pro Leu Gln Gly Ser Phe Asp Val Ala His 50 55 60

His Gly Gln Val Thr His Leu His Gly Arg Lys Ser Val Phe Asp Gly 65 70 75 80

Pro Thr Asp Val Leu Tyr Leu Pro Thr Gly Gln Thr Ala Thr Leu Ser 85 90 95

Gly Gln Gly Arg Val Ala Val Ala Glu Ala Pro Thr Gln Glu Pro Lys 100 105 110

Glu Trp Lys Tyr Ile Ala Pro Ala Glu Thr Pro Val Glu Leu Arg Gly 115 120 125

Ala Gly Arg Ser Ser Arg Gln Val His Asn Phe Gly Thr Pro Glu Ala 130 135 140

Leu Asp Ala Ala Arg Leu Ile Val Cys Glu Val Ile Thr Pro Gly Glu 145 150 155 160

Asn Trp Ser Ser Tyr Pro Pro His Lys His Asp Glu His Ile Pro Gly
165 170 175

His Glu Ser Lys Leu Glu Glu Ile Tyr Tyr Phe Glu Ser Ala Pro Ser 180 185 190

Arg Val Gly Gly Arg Ala Glu Ala Ala Glu Gly Ala Phe Gly Met Phe 195 200 205

Ser Thr Tyr Ser Ser Pro Ala Gly Glu Ile Asp Ile Asn Ala Met Val 210 215 220

Tyr Ser Gly Asp Ile Ala Leu Val Pro Phe Gly Tyr His Gly Pro Ala 225 230 235 240

Val Ala Ala Pro Gly Tyr Asp Leu Tyr Tyr Leu Asn Val Met Ala Gly 245 250 255

Pro Asp Pro Glu Arg Ile Trp Leu Ile Asn Asp Asp Pro Ala His Ala 260 265 270

Trp Val Arg Asp Thr Trp Thr Gly Gln Ala Phe Asp Asp Arg Leu Pro 275 280 285

Tyr Glu Asn Ala Asn Lys Glu Gly 290 295

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Ala Glu Ala Ala Glu Gly Ala Phe Gly Met Phe Ser Thr Tyr Ser Ser

205 200 210 787 cca qcq qqq gag atc gat atc aac gcc atg gtg tac agc ggc gat atc Pro Ala Gly Glu Ile Asp Ile Asn Ala Met Val Tyr Ser Gly Asp Ile 220 qcq cta qtt cct ttc qqa tac cac ggc cct gcc gtg gca gca cct ggc 835 Ala Leu Val Pro Phe Gly Tyr His Gly Pro Ala Val Ala Ala Pro Gly 235 883 tat gac ttg tac tac ctc aac gtc atg gca gga cct gat ccg gag aga Tyr Asp Leu Tyr Tyr Leu Asn Val Met Ala Gly Pro Asp Pro Glu Arg atc tgg ctg att aac gat gac cca gcg cac gcc tgg gtt cga gat aca 931 Ile Trp Leu Ile Asn Asp Asp Pro Ala His Ala Trp Val Arg Asp Thr 964 tgg acc ggg caa gca ttt gat gat cgc ttg cca Trp Thr Gly Gln Ala Phe Asp Asp Arg Leu Pro <210> 156 <211> 288 <212> PRT <213> Corynebacterium glutamicum Met Arg Trp Phe His Lys Lys Gly Glu Leu Ala Arg Asp Gly Trp Gln 5 1.0 Ser Val Val Asp Ala Thr Thr Pro Gly Trp Glu Tyr Thr Gly Ile Arg Ile Ala Glu Leu Gly Ser Gly Glu Ser Leu Glu Leu Asn Asp Thr Gly 35 Val Glu Arg Ile Phe Ile Pro Leu Gln Gly Ser Phe Asp Val Ala His His Gly Gln Val Thr His Leu His Gly Arg Lys Ser Val Phe Asp Gly Pro Thr Asp Val Leu Tyr Leu Pro Thr Gly Gln Thr Ala Thr Leu Ser Gly Gln Gly Arg Val Ala Val Ala Glu Ala Pro Thr Gln Glu Pro Lys Glu Trp Lys Tyr Ile Ala Pro Ala Glu Thr Pro Val Glu Leu Arg Gly 120 Ala Gly Arg Ser Ser Arg Gln Val His Asn Phe Gly Thr Pro Glu Ala 130 135 Leu Asp Ala Ala Arg Leu Ile Val Cys Glu Val Ile Thr Pro Gly Glu 155 Asn Trp Ser Ser Tyr Pro Pro His Lys His Asp Glu His Ile Pro Gly 165 170

His Glu Ser Lys Leu Glu Glu Ile Tyr Tyr Phe Glu Ser Ala Pro Ser 180

Arg Val Gly 195

Ser Thr Tyr Ser Ser Pro Ala Gly 200

Tyr Ser Gly Asp Ile Ala Leu 215

Val Ala Ala Ala Pro Gly Tyr Asp Leu Tyr Tyr 250

Fro Asp Pro Glu Arg Ile Trp Leu 265

Trp Val Arg 275

Arg Ser In Trp Val Arg Asp Thr Trp Thr Gly 280

Fro Asp Pro Asp Pro 275

Fro Asp Pro Asp Thr Trp Thr Gly 280

Fro Ala Phe Gly Tyr Asp Leu Tyr Tyr Leu Asp Asp Asp Pro Ala 270

Fro Asp Pro 260

Fro Asp Pro 275

Fro Asp Pro 285

Fro Asp Asp Asp Pro 285

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236

75 80 85 70 373 gga cat atg acg tgg gat Gly His Met Thr Trp Asp <210> 158 <211> 91 <212> PRT <213> Corynebacterium glutamicum <400> 158 Met Gln Lys Asn Ile Leu Lys Ser Gly Ile Glu Ile Ser Glu Leu Gly Leu Gly Cys Met Ser Leu Gly Thr Asp Tyr Lys Lys Ala Gln Pro Ile Ile Glu Ser Ala Ile Asp Asn Gly Ile Thr Tyr Phe Asp Thr Ala Asp Ile Tyr Asp Gln Gly Val Asn Glu Glu Ile Val Gly Lys Ala Leu Lys Lys Tyr Gln Asn Arg Asp Asp Ile Val Ile Gly Thr Lys Val Gly Asn 70 Arg Leu Thr Asp Asp Gly His Met Thr Trp Asp 85 <210> 159 <211> 376 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(376) <223> FRXA02829 <400> 159 tttttcgttt aatctcatat ttaaacacgt tccttttaat tggttttata aattgataaa 60 115 ctgaattcgt cagttaaagt gtatcgaaag gagactggac atg caa aaa aat att Met Gln Lys Asn Ile cta aaa agt ggc atc gaa att tct gaa ctt ggg tta ggt tgc atg agt 163 Leu Lys Ser Gly Ile Glu Ile Ser Glu Leu Gly Leu Gly Cys Met Ser 211 tta ggc aca gat tat aaa aaa gcg caa cca att att gaa agt gca att Leu Gly Thr Asp Tyr Lys Lys Ala Gln Pro Ile Ile Glu Ser Ala Ile 259 gat aat ggt att acg tat ttt gat act gca gat att tac gat caa gga Asp Asn Gly Ile Thr Tyr Phe Asp Thr Ala Asp Ile Tyr Asp Gln Gly

gtt aat gaa gaa att gtt ggt aaa gcc te Val Asn Glu Glu Ile Val Gly Lys Ala Lo 55 60	
gat gac atc gtt atc gga act aaa gtt ga Asp Asp Ile Val Ile Gly Thr Lys Val G 70	ga aat cga tta act gac gat 355 Ly Asn Arg Leu Thr Asp Asp 80 85
gga cat atg acg tgg gga tcc Gly His Met Thr Trp Gly Ser 90	376
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Ile Glu Ser Ala Ile Asp Asn Gly Ile T 35 40	nr Tyr Phe Asp Thr Ala Asp 45
Ile Tyr Asp Gln Gly Val Asn Glu Glu I 50 55	le Val Gly Lys Ala Leu Lys 60
Lys Tyr Gln Asn Arg Asp Asp Ile Val I 65 70	le Gly Thr Lys Val Gly Asn 75 80
Arg Leu Thr Asp Asp Gly His Met Thr T 85	rp Gly Ser 90
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tca tat ctg acc gac atg gac ggc gtc c Ser Tyr Leu Thr Asp Met Asp Gly Val L 10	
att ccg ggt gca gat cgt ttt ctt cag t Ile Pro Gly Ala Asp Arg Phe Leu Gln S	

25 30 35

						aac Asn										259
						tcc Ser 60										307
						act Thr										355
						gtt Val										403
						ttg Leu										451
						tat Tyr										499
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act Thr 150	gga Gly	cct Pro	tca Ser	cca Pro	agt Ser 155	ggc Gly	att Ile	ttg Leu	cct Pro	gct Ala 160	act Thr	ggc Gly	tct Ser	gtc Val	gcc Ala 165	595
gca Ala	ctt Leu	att Ile	acc Thr	gca Ala 170	gct Ala	act Thr	ggc Gly	gct Ala	gag Glu 175	cct Pro	tat Tyr	tac Tyr	atc Ile	ggc Gly 180	aag Lys	643
cca Pro	aac Asn	cct Pro	gtg Val 185	atg Met	atg Met	cgc Arg	agt Ser	gcg Ala 190	ctg Leu	aac Asn	acc Thr	atc Ile	ggg Gly 195	gcg Ala	cat His	691
	, ,			_	_	atc Ile		_	_	-	-		-			739
						ctg Leu 220										787
						cgc Arg										835
						ctt Leu										883
						cca Pro										925

tagtattctg taggtcatgg cat

948

<210> 162

<211> 275

<212> PRT

<213> Corynebacterium glutamicum

<400> 162

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Thr Asp Asn Asn Val Glu Phe Met Val Leu Thr Asn Asn Ser Ile Phe 35 40 45

Thr Pro Arg Asp Leu Ser Ala Arg Leu Lys Thr Ser Gly Leu Asp Ile 50 60

Pro Pro Glu Arg Ile Trp Thr Ser Ala Thr Ala Thr Ala His Phe Leu 65 70 75 80

Lys Ser Gln Val Lys Glu Gly Thr Ala Tyr Val Val Gly Glu Ser Gly 85 90 95

Leu Thr Thr Ala Leu His Thr Ala Gly Trp Ile Leu Thr Asp Ala Asn 100 105 110

Pro Glu Phe Val Val Leu Gly Glu Thr Arg Thr Tyr Ser Phe Glu Ala 115 120 125

Ile Thr Thr Ala Ile Asn Leu Ile Leu Gly Gly Ala Arg Phe Ile Cys 130 140

Thr Asn Pro Asp Val Thr Gly Pro Ser Pro Ser Gly Ile Leu Pro Ala 145 150 155 160

Thr Gly Ser Val Ala Ala Leu Ile Thr Ala Ala Thr Gly Ala Glu Pro 165 170 175

Tyr Tyr Ile Gly Lys Pro Asn Pro Val Met Met Arg Ser Ala Leu Asn 180 185 190

Thr Ile Gly Ala His Ser Glu His Thr Val Met Ile Gly Asp Arg Met 195 200 205

Asp Thr Asp Val Lys Ser Gly Leu Glu Ala Gly Leu Ser Thr Val Leu 210 215 220

Val Arg Ser Gly Ile Ser Asp Asp Ala Glu Ile Arg Arg Tyr Pro Phe 225 230 235 240

Arg Pro Thr His Val Ile Asn Ser Ile Ala Asp Leu Ala Asp Cys Trp
245 250 255

Asp Asp Pro Phe Gly Asp Gly Ala Phe His Val Pro Asp Glu Gln Gln 260 265 270

Phe Thr Asp 275

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					gct Ala											643
					atg Met											691
					atg Met											739
					ggc Gly											787
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					gat Asp											883
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Ile Thr Thr Ala Ile Asn Leu Ile Leu Gly Gly Ala Arg Phe Ile Cys Thr Asn Pro Asp Val Thr Gly Pro Ser Pro Ser Gly Ile Leu Pro Ala 155 Thr Gly Ser Val Ala Ala Leu Ile Thr Ala Ala Thr Gly Ala Glu Pro Tyr Tyr Ile Gly Lys Pro Asn Pro Val Met Met Arg Ser Ala Leu Asn 180 185 190 Thr Ile Gly Ala His Ser Glu His Thr Val Met Ile Gly Asp Arg Met 200 Asp Thr Asp Val Lys Ser Gly Leu Glu Ala Gly Leu Ser Thr Val Leu 215 Val Arg Ser Gly Ile Ser Asp Asp Ala Glu Ile Arg Arg Tyr Pro Phe Arg Pro Thr His Val Ile Asn Ser Ile Ala Asp Leu Ala Asp Cys Trp 255 245 Asp Asp Pro Phe Gly Asp Gly Ala Phe His Val Pro Asp Glu Gln Gln Phe Thr Asp 275 <210> 165 <211> 1128 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(1105) <223> RXA00794 <400> 165 gcqqqttqat acaqcccaaq cqccqataca tttataatqc qcctagatac gtgcaaccca 60 cgtaaccagg tcagatcaag tgccccagga ggcccttcag atg aac cta aag aac Met Asn Leu Lys Asn ccc gaa acg cca gac cgt aac ctt gct atg gag ctg gtg cga gtt acg Pro Glu Thr Pro Asp Arg Asn Leu Ala Met Glu Leu Val Arg Val Thr gaa gca gct gca ctg gct tct gga cgt tgg gtt gga cgt ggc atg aag 211 Glu Ala Ala Ala Leu Ala Ser Gly Arg Trp Val Gly Arg Gly Met Lys aat gaa ggc gac ggt gcc gct gtt gac gcc atg cgc cag ctc atc aac 259 Asn Glu Gly Asp Gly Ala Ala Val Asp Ala Met Arg Gln Leu Ile Asn tca qtg acc atg aag qqc qtc qtt qtt atc qqc gag qgc gaa aaa gac 307

Ser	Val 55	Thr	Met	Lys	Gly	Val 60	Val	Val	Ile	Gly	Glu 65	Gly	Glu	Lys	Asp	
							ggc Gly									355
cct Pro	gag Glu	gtt Val	gat Asp	atc Ile 90	gca Ala	gtt Val	gac Asp	cca Pro	gtt Val 95	gac Asp	ggc Gly	acc Thr	acc Thr	ctg Leu 100	atg Met	403
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							tcc Ser 125									499
							aag Lys									547
							aag Lys									595
-		-	-			•	cgt Arg		-			_	_		-	643
							aag Lys									691
_	-		-	-	-	-	gct Ala 205	_	_					-		739
							cca Pro									787
							atc Ile									835
_			-	_	_	_	cac His	-	_		_	-		_	_	883
							gtg Val									931
-							gac Asp 285	-		-		-			-	979
-			-			_	tcc Ser	_	-	_	-	_	_			1027

295 300 305

acc atc cgc cac atc gag tct gtc cac cag ctg tcc aag ctg cag gaa 1075
Thr Ile Arg His Ile Glu Ser Val His Gln Leu Ser Lys Leu Gln Glu
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Gly Arg Gly Met Lys Asn Glu Gly Asp Gly Ala Ala Val Asp Ala Met 35 40 45

Arg Gln Leu Ile Asn Ser Val Thr Met Lys Gly Val Val Val Ile Gly 50 . 60

Glu Gly Glu Lys Asp Glu Ala Pro Met Leu Tyr Asn Gly Glu Glu Val 65 70 75 80

Gly Thr Gly Phe Gly Pro Glu Val Asp Ile Ala Val Asp Pro Val Asp 85 90 95

Gly Thr Thr Leu Met Ala Glu Gly Arg Pro Asn Ala Ile Ser Ile Leu 100 105 110

Ala Ala Ala Glu Arg Gly Thr Met Tyr Asp Pro Ser Ser Val Phe Tyr 115 120 125

Met Lys Lys Ile Ala Val Gly Pro Glu Ala Ala Gly Lys Ile Asp Ile 130 135 140

Glu Ala Pro Val Ala His Asn Ile Asn Ala Val Ala Lys Ser Lys Gly 145 150 155 160

Ile Asn Pro Ser Asp Val Thr Val Val Leu Asp Arg Pro Arg His
165 170 175

Ile Glu Leu Ile Ala Asp Ile Arg Arg Ala Gly Ala Lys Val Arg Leu 180 185 190

Ile Ser Asp Gly Asp Val Ala Gly Ala Val Ala Ala Ala Gln Asp Ser 195 200 205

Asn Ser Val Asp Ile Met Met Gly Thr Gly Gly Thr Pro Glu Gly Ile 210 215 220

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Leu Ala Pro Met	Asn Asp Phe 245	e Glu Arg Gln Ly: 250	s Ala His Asp Ala Gly 255	
Leu Val Leu Asp 260	Gln Val Leu	His Thr Asn Asp 265	p Leu Val Ser Ser Asp 270	
Asn Cys Tyr Phe 275	Val Ala Thi	r Gly Val Thr Asi 280	n Gly Asp Met Leu Arg 285	
Gly Val Ser Tyr 290	Arg Ala Asr 295		r Arg Ser Leu Val Met 300	
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tgcatgcaga ttat cgggtggcgt cgaa tat ccg cat ttg	aagcat tttta tgg gag tco	aaagga gtttaagac	g atg aag ttt gtt atg Met Lys Phe Val Met	-
tgcatgcaga ttat cgggtggcgt cgaa  tat ccg cat ttg Tyr Pro His Leu cat gag cgg gtt	tgg gag tcc Trp Glu Ser 10 gag gat att	aaagga gtttaagac c acg acc gct gto r Thr Thr Ala Va 15 c aaa gat gca gao	g atg aag ttt gtt atg Met Lys Phe Val Met 1 5 c att gag ggt ggc gga l Ile Glu Gly Gly	115
tgcatgcaga ttat cgggtggcgt cgaa  tat ccg cat ttg Tyr Pro His Leu  cat gag cgg gtt His Glu Arg Val 25 ggt tca gcg ccg	tgg gag tcc Trp Glu Ser 10 gag gat att Glu Asp Ile	aaagga gtttaagace c acg acc gct gte c Thr Thr Ala Vai 15 c aaa gat gca gae c Lys Asp Ala Asp 30 g gat ttg ccg gae	g atg aag ttt gtt atg Met Lys Phe Val Met 1 5 c att gag ggt ggc gga l Ile Glu Gly Gly Gly 20 c ttc att ttc ttt aat	115
tgcatgcaga ttat cgggtggcgt cgaa  tat ccg cat ttg Tyr Pro His Leu  cat gag cgg gtt His Glu Arg Val	tgg gag tcc Trp Glu Ser 10 gag gat att Glu Asp Ile gag ttc ccc Glu Phe Pro	aaagga gtttaagace c acg acc gct gte c Thr Thr Ala Vai 15 c aaa gat gca gae c Lys Asp Ala Asp 30 g gat ttg ccg gae o Asp Leu Pro Glu 45 c gat gcg ctg gte c Asp Ala Leu Vai	g atg aag ttt gtt atg Met Lys Phe Val Met 1 5  c att gag ggt ggc gga 1 Ile Glu Gly Gly Gly 20  c ttc att ttc ttt aat Phe Ile Phe Phe Asn 35  g aac atc aag ttc gtg 1 Asn Ile Lys Phe Val	115 163 211
tgcatgcaga ttat cgggtggcgt cgaa  tat ccg cat ttg Tyr Pro His Leu  cat gag cgg gtt His Glu Arg Val	tgg gag tcc Trp Glu Ser 10  gag gat att Glu Asp Ile  gag ttc ccc Glu Phe Pro  gcg ggt att Ala Gly Ile  cgt tgg gca	aaagga gtttaagace c acg acc gct gte c Thr Thr Ala Vai 15 c aaa gat gca gae c Lys Asp Ala Asp 30 g gat ttg ccg gae o Asp Leu Pro Glu 45 c gat gcg ctg gte c Asp Ala Leu Vai 0	g atg aag ttt gtt atg Met Lys Phe Val Met 1 5  att gag ggt ggc gga l Ile Glu Gly Gly Gly 20  c ttc att ttc ttt aat Phe Ile Phe Phe Asn 35  g aac atc aag ttc gtg l Asn Ile Lys Phe Val 50  c aag cgt ggt gtc gtc l Lys Arg Gly Val Val 65  c ctg tac gct gac acc y Leu Tyr Ala Asp Thr	115 163 211 259

		90			95				100		
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			cat His								499
			cgt Arg 140								547
			aat Asn								595
			gat Asp								643
			ctg Leu								691
			ggc Gly								739
			ctg Leu 220								787
			gcg Ala								835
			cac His								883
			aac Asn								931
			aac Asn								979
			gtg Val 300			tago	gcctt	itt a	atggt	gtgat	1032
ccg											1035

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<sup>&</sup>lt;211> 304

<sup>&</sup>lt;212> PRT

<sup>&</sup>lt;213> Corynebacterium glutamicum

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gag tcc acc att ggt tta att ctg gcg cag atg cac atg cat gcg Glu Ser Thr Ile Gly Leu Ile Leu Ala Gln Met His Met His Ala 20 25 30	acg 153 Thr
act cgt ttg gct aag tcg tgg agc gtg cgg cct gag gtg gaa aac Thr Arg Leu Ala Lys Ser Trp Ser Val Arg Pro Glu Val Glu Asn 35 40 45	
aag toa tgg ctg cat gac aat aaa act gtc gct att ttg ggc gcc Lys Ser Trp Leu His Asp Asn Lys Thr Val Ala Ile Leu Gly Ala 55 : 60 65	
ggc att ggc gtg cgt ctg ctg gaa atg ctc aag ccg ttc aac gtg Gly Ile Gly Val Arg Leu Leu Glu Met Leu Lys Pro Phe Asn Val 70 75 80	
acc att gcg gtt aat aac tct ggt cgt ccg gtg gaa ggt gca gat Thr Ile Ala Val Asn Asn Ser Gly Arg Pro Val Glu Gly Ala Asp 85 90 95	
acc ttc gcc atg gat aag gct gag cac gtg tgg gct gag gct gat Thr Phe Ala Met Asp Lys Ala Glu His Val Trp Ala Glu Ala Asp 100 105 110	
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gca gaa act ttg ggc aag atg aag cct tct gcc gtg gtg gtc aat Ala Glu Thr Leu Gly Lys Met Lys Pro Ser Ala Val Val Val Asn 135 140	
ggg cgt ggc ccg ctg atc aac acc gat gat ctg gtg gat gca ttg Gly Arg Gly Pro Leu Ile Asn Thr Asp Asp Leu Val Asp Ala Leu 150 155 160	
aac ggc acc att gcg ggt gct gcg ctg gac gtt acc gat cct gag Asn Gly Thr Ile Ala Gly Ala Ala Leu Asp Val Thr Asp Pro Glu 165 170 175	

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						aat Asn										681
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ccg																779
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	0> 1		** 1		_	* 1 -		<b>3</b> 3.	7.1 -	<b>61</b>	<b>T</b>	m	21-	3	m)	
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Val	Ala	Glu	Ser 20	Thr	Ile	Gly	Leu	Ile 25	Leu	Ala	Gln	Met	His 30	Met	His	
Ala	Thr	Thr 35	Arg	Leu	Ala	Lys	Ser 40	Trp	Ser	Val	Arg	Pro 45	Glu	Val	Glu	
Asn	Asn 50	Lys	Ser	Trp	Leu	His 55	Asp	Asn	Lys	Thr	Val 60	Ala	Ile	Leu	Gly	
Ala 65	Gly	Gly	Ile	Gly	Val 70	Arg	Leu	Leu	Glu	Met 75	Leu	Lys	Pro	Phe	Asn 80	
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Leu	Asn	Asn	Gly	Thr 165	Ile	Ala	Gly	Ala	Ala 170	Leu	Asp	Val	Thr	Asp 175	Pro	
Glu	Pro	Leu	Pro 180	Asp	Ser	His	Pro	Leu 185	Trp	Glu	Met	Asp	Asn 190	Val	Val	

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135 140 145 atc atg gac agc cac gac ctt gac gac gac ggc gac gtt gcc gtt gtc 595 Ile Met Asp Ser His Asp Leu Asp Asp Asp Arg Asp Val Ala Val Val age cae gge gee ate ege ate gtg gea aca cae gea act ggt gtg 643 Ser His Gly Ala Val Ile Arg Ile Val Ala Thr His Ala Thr Gly Val gat ecc aac ttt geg tte aac acc tac etg gge aac tge ege tte gtg 691 Asp Pro Asn Phe Ala Phe Asn Thr Tyr Leu Gly Asn Cys Arg Phe Val 739 gtg ctg gag cca aac ggt aag aaa ttc agc caa tgg gat gtt gtg cgc Val Leu Glu Pro Asn Gly Lys Lys Phe Ser Gln Trp Asp Val Val Arg 205 789 tgg act gac agc cca ctg cca tgg cag gag taattgagac caaaggctcg Trp Thr Asp Ser Pro Leu Pro Trp Gln Glu 792 gat <210> 172 <211> 223 <212> PRT <213> Corynebacterium glutamicum <400> 172 Met Ala Gly Arg Ile Ile Leu Leu Arg His Gly Gln Thr His Asn Asn Val Lys His Leu Leu Asp Thr Arg Pro Pro Gly Ala Glu Leu Thr Asp Leu Gly Arg Lys Gln Ala Leu Glu Val Gly His Glu Leu Ala Thr Tyr 40 Ser Gly Glu Arg Leu Ala His Val Tyr Ser Ser Ile Val Leu Arg Ala Gln Gln Thr Ala Val Leu Ala Thr Ser Thr Phe Glu Lys Ala Arg Asp Met Gln Ser Gly Ala Ile Pro Leu Asp Val Val Glu Gly Ile Gln Glu Ile Asn Val Gly Asp Phe Glu Met Arg Gly Asp Glu Glu Ala His Met 100 Asn Tyr Ser Arg Ala Leu Asn Gly Trp Leu His Gly Asp Pro Ala Ala Gly Leu Pro Gly Gly Glu Thr Tyr Lys Asp Val Leu Asn Arg Tyr Gln Pro Thr Leu Asp Arg Ile Met Asp Ser His Asp Leu Asp Asp Asp Arg 145

Asp Val Ala Val Val Ser His Gly Ala Val Ile Arg Ile Val Ala Thr His Ala Thr Gly Val Asp Pro Asn Phe Ala Phe Asn Thr Tyr Leu Gly 185 Asn Cys Arg Phe Val Val Leu Glu Pro Asn Gly Lys Lys Phe Ser Gln 200 Trp Asp Val Val Arg Trp Thr Asp Ser Pro Leu Pro Trp Gln Glu 215 <210> 173 <211> 336 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(313) <223> RXN03087 <400> 173 gttgccgcca gccgttccag ggcgcttgag ctggtcagcg acatcgcaat gatcaaccag 60 gaatacctgg aaaagagctg atattgatag ggtttaagtc atg aag atc tac gca Met Lys Ile Tyr Ala 1 163 cct ttt gct gga atc gtc cac tat ttt gtc gat gaa ggc gat ccc gtg Pro Phe Ala Gly Ile Val His Tyr Phe Val Asp Glu Gly Asp Pro Val 20 10 qaa acc ggc atg caa ctg gga acg gta gaa acc atc aaa ctc gag gca 211 Glu Thr Gly Met Gln Leu Gly Thr Val Glu Thr Ile Lys Leu Glu Ala 35 25 30 cca atc atg gca ccg gga cct ggc atc gta gct aag gtt tct ttt gat 259 Pro Ile Met Ala Pro Gly Pro Gly Ile Val Ala Lys Val Ser Phe Asp 40 45 gat ttc tcc gac gtc acc ggc ggc gat gaa ctc ctc gaa ttg gag gca 307 Asp Phe Ser Asp Val Thr Gly Gly Asp Glu Leu Leu Glu Leu Glu Ala 55 60 336 aag aac taatgggtca aacccgcatc att Lys Asn 70 <210> 174 <211> 71 <212> PRT <213> Corynebacterium glutamicum <400> 174 Met Lys Ile Tyr Ala Pro Phe Ala Gly Ile Val His Tyr Phe Val Asp 15 Glu Gly Asp Pro Val Glu Thr Gly Met Gln Leu Gly Thr Val Glu Thr

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25

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Ser Ile Val Val Ala Val Leu Asn Ser Leu Ala Arg Glu Gly Lys Ile Asp Val Ser Val Ala Ala Gln Ala Ala Glu Lys Phe Lys Leu Asp Asp Pro Thr Ser Val Ser Val Asp Pro Asn Ala Pro Glu Glu 85 <210> 179 <211> 1953 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(1930) <223> RXN02591 <400> 179 atgtgtccgt tgtctcacct aaagttttaa ctagttctgt atctgaaagc tacgctaggg 60 ggcgagaact ctgtcgaatg acacaaaatc tggagaagta atg act act gct gca Met Thr Thr Ala Ala 1 163 atc agg ggc ctt cag ggc gag gcg ccg acc aag aat aag gaa ctg ctg Ile Arg Gly Leu Gln Gly Glu Ala Pro Thr Lys Asn Lys Glu Leu Leu 15 10 aac tgg atc gca gac gcc gtc gag ctc ttc cag cct gag gct gtt gtg 211 Asn Trp Ile Ala Asp Ala Val Glu Leu Phe Gln Pro Glu Ala Val Val 25 30 ttc gtt gat gga tcc cag gct gag tgg gat cgc atg gcg gag gat ctt 259 Phe Val Asp Gly Ser Gln Ala Glu Trp Asp Arg Met Ala Glu Asp Leu 40 gtt qaa gcc ggt acc ctc atc aag ctc aac gag gaa aag cgt ccg aac 307 Val Glu Ala Gly Thr Leu Ile Lys Leu Asn Glu Glu Lys Arg Pro Asn 55 355 age tac cta get egt tee aac cea tet gae gtt geg ege gtt gag tee Ser Tyr Leu Ala Arg Ser Asn Pro Ser Asp Val Ala Arg Val Glu Ser 70 75 cgc acc ttc atc tgc tcc gag aag gaa gaa gat gct ggc cca acc aac 403 Arg Thr Phe Ile Cys Ser Glu Lys Glu Glu Asp Ala Gly Pro Thr Asn 90 100 aac tgg gct cca cca cag gca atg aag gac gaa atg tcc aag cat tac 451 Asn Trp Ala Pro Pro Gln Ala Met Lys Asp Glu Met Ser Lys His Tyr 105 110 115 499 gct ggt tcc atg aag ggg cgc acc atg tac gtc gtg cct ttc tgc atg Ala Gly Ser Met Lys Gly Arg Thr Met Tyr Val Val Pro Phe Cys Met 120 125

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tcc gag ta Ser Glu Ty 150	c gtt gtc r Val Val	atg tcc Met Ser 155	atg cgc Met Arc	atc J Ile	atg a Met 1	acc co Thr Ar	gc atg gg Met	ggt Gly	att Ile 165	595
gaa gcg ct Glu Ala Le	g gac aag u Asp Lys 170	atc ggc Ile Gly	gcg aad Ala Asi	ggc Gly 175	agc ( Ser )	ttc gt Phe Va	c agg al Arg	tgc Cys 180	ctc Leu	643
cac tcc gt His Ser Va	t ggt gct 1 Gly Ala 185			Gly						691
	c gac acc n Asp Thr						u Thr			739
att tgg to Ile Trp Se 215	c tac ggt r Tyr Gly	tcc ggc Ser Gly 220	tac ggo Tyr Gl	gga Gly	Asn A	gca at Ala IJ 225	c ctg le Leu	gca Ala	aag Lys	787
	c gca ctg r Ala Leu									835
	t gag cac a Glu His 250									883
	c cac atc r His Ile 265			Pro						931
	cc atg atc a Met Ile 80						r Ala			979
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gca gtt aa Ala Val As 310	ac cca gaa sn Pro Glu	aat ggt Asn Gly 315	ttc ttc Phe Pho	ggt Gly	gtt ( Val 2 320	get ed Ala Pr	ca ggc co Gly	acc Thr	aac Asn 325	1075
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ctg ttc ac Leu Phe Th	cc aac gtg or Asn Val 345	gca ctc Ala Leu	acc gad Thr Asj 35	Asp	ggc (	gac at Asp Il	tc tgg le Trp 355	tgg Trp	gaa Glu	1171
	ac ggc gac sp Gly Asp 50						p Met			1219

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gac tgg gaa ggc Asp Trp Glu Gly	gtc aag atc Val Lys Ile 410	gac gca atc Asp Ala Ile 415	ctc ttc ggt Leu Phe Gly	gga cgt cgc Gly Arg Arg 420	1363
gca gac acc gtc Ala Asp Thr Val 425	cca ctg gtt Pro Leu Val	acc cag acc Thr Gln Thr 430	tac gac tgg Tyr Asp Trp	gag cac ggc Glu His Gly 435	1411
acc atg gtt ggt Thr Met Val Gly 440	gca ctg ctc Ala Leu Leu	gca tcc ggt Ala Ser Gly 445	cag acc gca Gln Thr Ala 450	gct tcc gca Ala Ser Ala	1459
gaa gca aag gtc Glu Ala Lys Val 455					1507
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ggt aac aag ggt Gly Asn Lys Gly					1603
ttc cgc cgt ggc Phe Arg Arg Gly 505					1651
aac tct cgc gtt Asn Ser Arg Val 520	ctg aag tgg Leu Lys Trp	gtc atc gac Val Ile Asp 525	cgc atc gaa Arg Ile Glu 530	ggc cac gtt Gly His Val	1699
ggc gca gac gag Gly Ala Asp Glu 535					1747
gac ctc gac ggc Asp Leu Asp Gly 550					1795
acc gct cct gca Thr Ala Pro Ala					1843
tac ctc act ttc Tyr Leu Thr Phe 585					1891
ttc gat gct ctg Phe Asp Ala Leu 600				taaagttcac	1940
gcttaagaac tgc					1953

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Val Pro Phe Cys Met Gly Pro Ile Ser Asp Pro Asp Pro Lys Leu Gly

Val Gln Leu Thr Asp Ser Glu Tyr Val Val Met Ser Met Arg Ile Met

Thr Arg Met Gly Ile Glu Ala Leu Asp Lys Ile Gly Ala Asn Gly Ser

Phe Val Arg Cys Leu His Ser Val Gly Ala Pro Leu Glu Pro Gly Gln

Glu Asp Val Ala Trp Pro Cys Asn Asp Thr Lys Tyr Ile Thr Gln Phe

280

155

135

150

275

Pro Glu Thr Lys Glu Ile Trp Ser Tyr Gly Ser Gly Tyr Gly Gly Asn 210

Ala Ile Leu Ala Lys Lys Cys Tyr Ala Leu Arg Ile Ala Ser Val Met 225

Ala Arg Glu Glu Gly Trp Met Ala Glu His Met Leu Ile Leu Lys Leu 255

Ile Asn Pro Glu Gly Lys Ala Tyr His Ile Ala Ala Ala Phe Pro Ser 260

Ala Cys Gly Lys Thr Asn Leu Ala Met Ile Thr Pro Thr Ile Pro Gly

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Glu 305	Asp	Gly	Leu	Tyr	Ala 310	Val	Asn	Pro	Glu	Asn 315	Gly	Phe	Phe	Gly	Val 320
Ala	Pro	Gly	Thr	Asn 325	Tyr	Ala	Ser	Asn	Pro 330	Ile	Ala	Met	Lys	Thr 335	Met
Glu	Pro	Gly	Asn 340	Thr	Leu	Phe	Thr	Asn 345	Val	Ala	Leu	Thr	Asp 350	Asp	Gly
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His 385	Pro	Asn	Ser	Arg	Tyr 390	Cys	Val	Ala	Ile	Asp 395	Gln	Ser	Pro	Ala	Ala 400
Ala	Pro	Glu	Phe	Asn 405	Asp	Trp	Glu	Gly	Val 410	Lys	Ile	Asp	Ala	Ile 415	Leu
Phe	Gly	Gly	Arg 420	Arg	Ala	Asp	Thr	Val 425	Pro	Leu	Val	Thr	Gln 430	Thr	Tyr
Asp	Trp	Glu 435	His	Gly	Thr	Met	Val 440	Gly	Ala	Leu	Leu	Ala 445	Ser	Gly	Gln
Thr	Ala 450	Ala	Ser	Ala	Glu	Ala 455	Lys	Val	Gly	Thr	Leu 460	Arg	His	Asp	Pro
Met 465	Ala	Met	Leu	Pro	Phe 470	Ile	Gly	Tyr	Asn	Ala 475	Gly	Glu	Tyr	Leu	Gln 480
Asn	Trp	Ile	Asp	Met 485	Gly	Asn	Lys	Gly	Gly 490	Asp	Lys	Met	Pro	Ser 495	Ile
Phe	Leu	Val	Asn 500	Trp	Phe	Arg	Arg	Gly 505	Glu	Asp	Gly	Arg	Phe 510	Leu	Trp
Pro	Gly	Phe 515	Gly	Asp	Asn	Ser	Arg 520	Val	Leu	Lys	Trp	Val 525	Ile	Asp	Arg
Ile	Glu 530	Gly	His	Val	Gly	Ala 535	Asp	Glu	Thr	Val	Val 540	Gly	His	Thr	Ala
Lys 545	Ala	Glu	Asp	Leu	Asp 550	Leu	Asp	Gly	Leu	Asp 555	Thr	Pro	Ile	Glu	Asp 560
Val	Lys	Glu	Ala	Leu 565	Thr	Ala	Pro	Ala	Glu 570	Gln	Trp	Ala	Asn	Asp 575	Val
Glu	Asp	Asn	Ala 580	Glu	Tyr	Leu	Thr	Phe 585	Leu	Gly	Pro	Arg	Val 590	Pro	Ala
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cgt Arg	cca Pro	cgc Arg 200	gtt Val	gag Glu	gga Gly	ttt Phe	ggt Gly 205	ctt Leu	gaa Glu	aac Asn	act Thr	ggc Gly 210	gtt Val	aag Lys	ctc Leu	739
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						gtt Val										1075
			Ala			gaa Glu				Leu					Leu	1123
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Ile Ile Ala Thr Gly Ser Val Val Asn Thr Leu Arg Gly Val Asp Phe 65 70 75 80

Ser Glu Asn Val Val Ser Phe Glu Glu Gln Ile Leu Asn Pro Val Ala 85 90 95

Pro Lys Lys Met Val Ile Val Gly Ala Gly Ala Ile Gly Met Glu Phe 100 105 110

Ala Tyr Val Leu Gly Asn Tyr Gly Val Asp Val Thr Val Ile Glu Phe 115 120 125

Met Asp Arg Val Leu Pro Asn Glu Asp Ala Glu Val Ser Lys Val Ile 130 135 140

Ala Lys Ala Tyr Lys Lys Met Gly Val Lys Leu Leu Pro Gly His Ala 145 150 155 160

Thr Thr Ala Val Arg Asp Asn Gly Asp Phe Val Glu Val Asp Tyr Gln 165 170 175

Lys Lys Gly Ser Asp Lys Thr Glu Thr Leu Thr Val Asp Arg Val Met 180 185 190

Val Ser Val Gly Phe Arg Pro Arg Val Glu Gly Phe Gly Leu Glu Asn 195 200 205

Thr Gly Val Lys Leu Thr Glu Arg Gly Ala Ile Glu Ile Asp Asp Tyr 210 215 220

Met Arg Thr Asn Val Asp Gly Ile Tyr Ala Ile Gly Asp Val Thr Ala 225 230 235 240

Lys Leu Gln Leu Ala His Val Ala Glu Ala Gln Gly Ile Val Ala Ala 245 250 255

Glu Thr Ile Ala Gly Ala Glu Thr Gln Thr Leu Gly Asp Tyr Met Met 260 265 270

Met Pro Arg Ala Thr Phe Cys Asn Pro Gln Val Ser Ser Phe Gly Tyr 275 280 285

Thr Glu Glu Gln Ala Lys Glu Lys Trp Pro Asp Arg Glu Ile Lys Val

290 295 300 Ala Ser Phe Pro Phe Ser Ala Asn Gly Lys Ala Val Gly Leu Ala Glu 310 Thr Asp Gly Phe Ala Lys Ile Val Ala Asp Ala Glu Phe Gly Glu Leu Leu Gly Ala His Leu Val Gly Ala Asn Ala Ser Glu Leu Ile Asn Glu Leu Val Leu Ala Gln Asn Trp Asp Leu Thr Thr Glu Glu Ile Ser Arg 360 Ser Val His Ile His Pro Thr Leu Ser Glu Ala Val Lys Glu Ala Ala 375 380 His Gly Ile Ser Gly His Met Ile Asn Phe 390 <210> 183 <211> 294 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(271) <223> RXS01261 <400> 183 qtqqqtqttt ttcattttct tccactctaa aattaagtat qqaaaaccaa ccgcacccgg 60 atgcacgaca atgacccact aaacacgtat ccttgaatgc gtg act gaa cat tat Val Thr Glu His Tyr 1 gac gta gta gta ctc gga gcc ggc ccc ggt ggc tat gtc tcc gcc atc Asp Val Val Val Leu Gly Ala Gly Pro Gly Gly Tyr Val Ser Ala Ile 10 cgt gca gcg cag ctt ggc aag aag gtt gct gta att gag aag cag tac 211 Arg Ala Ala Gln Leu Gly Lys Lys Val Ala Val Ile Glu Lys Gln Tyr 25 tgg ggt ggt gtt tgc cta aac gtg ggc tgc att cct tcc aaa gtc tct 259 Trp Gly Gly Val Cys Leu Asn Val Gly Cys Ile Pro Ser Lys Val Ser 45 40 gat caa aaa cgc tgaagttgcc cataccttta ccc 294 Asp Gln Lys Arg 55 <210> 184 <211> 57 <212> PRT <213> Corynebacterium glutamicum <400> 184

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gcc gac ( Ala Asp													883
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cca gca Pro Ala													1219

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atq	agc	gaa	gaa	gag	cgc											1603
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Tyr Leu Pro Gly Ile Thr Leu Pro Glu Ser Leu Gln Val Thr Ser Ser 50 55 60

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Ser Gln Ala Leu Arg Gly Asn Leu Ala Glu Trp Lys Glu Thr Ile Pro 85 90 95

Gln Asp Ala Thr Leu Val Ser Leu Ala Lys Gly Ile Glu Lys Gly Thr 100 105 110

His Leu Arg Met Ser Glu Val Ile Ala Glu Val Thr Glu Ala Asp Pro 115 120 125

Ser Arg Ile Ala Val Leu Ser Gly Pro Asn Leu Ala Arg Glu Ile Ala 130 135 140

Glu Gly Gln Pro Ala Ala Thr Val Ile Ala Cys Pro Asp Glu Asn Arg 145 150 155 160

Ala Lys Leu Val Gln Ala Ala Val Ala Ala Pro Tyr Phe Arg Pro Tyr 165 170 175

Thr Asn Thr Asp Val Val Gly Thr Glu Ile Gly Gly Ala Cys Lys Asn 180 185 190

Val Ile Ala Leu Ala Cys Gly Ile Ser His Gly Tyr Gly Leu Gly Glu 195 200 205

Asn Thr Asn Ala Ser Leu Ile Thr Arg Gly Leu Ala Glu Ile Ala Arg 210 215 220

Leu Gly Ala Thr Leu Gly Ala Asp Ala Lys Thr Phe Ser Gly Leu Ala 225 230 235 240

Gly Met Gly Asp Leu Val Ala Thr Cys Ser Ser Pro Leu Ser Arg Asn 245 250 255

Arg Ser Phe Gly Glu Arg Leu Gly Gln Gly Glu Ser Leu Glu Lys Ala 260 265 270

Arg Glu Ala Thr Asn Gly Gln Val Ala Glu Gly Val Ile Ser Ser Gln 275 280 285

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Tyr Leu Pro 50	Gly Ile Thr	Leu Pro Glu 55	Ser Leu Gln 60	Val Thr Ser	Ser
Ala Thr Glu 65	Ala Leu Glu 70		Ile Val Val 75	Leu Ala Ile	Pro 80

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	cgt ggc Arg Gly 280		Gly L									979
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His Gly Gly Leu Arg Tyr Leu Glu Gln Tyr Asp Phe Gly Val Val Gln 65 70 75 80	
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Pro Glu Gly Leu Lys Gly Ala Trp Arg His Asp Asp Thr Leu Asn Leu 165 170 175	

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Ala Ala Val Asp Leu Ala Asp Val Leu Asp Arg Arg Ile Val Leu Gly Thr Leu Gly Tyr Val Gln Pro Ala Ala Val Arg Ala Thr Ala Glu Ala 520 Met Ala Gln Val Thr Gly Trp Ser Ala Glu Leu Ile Asp Ala Gln Cys 535 530 Gln Ser Tyr Leu Ala Lys Gln Asp Lys Ile Gln Ala Val Leu Lys Pro 550 . Tyr Arg <210> 193 <211> 900 <212> DNA <213> Corvnebacterium glutamicum <220> <221> CDS <222> (101)..(877) <223> RXA01242 <400> 193 cgccggcaac caaatgaggc ttttgggcgt tggacagtga gacaatgggt aagaaattcg 60 gacatattta gtaaattggc tttttgcttt aaggagtgac atg tac gca gag gag 115 Met Tyr Ala Glu Glu cgc cgt cga cag att gcc tca tta acg gca gtt gag gga cgt gta aat Arg Arg Arg Gln Ile Ala Ser Leu Thr Ala Val Glu Gly Arg Val Asn qtc aca gaa tta gcg ggc cga ttc gat gtc act gca gag acg att cga 211 Val Thr Glu Leu Ala Gly Arg Phe Asp Val Thr Ala Glu Thr Ile Arg 30 259 cga gac ctt gcg gtg cta gac cgc gag gga att gtt cac cgc gtt cac Arg Asp Leu Ala Val Leu Asp Arg Glu Gly Ile Val His Arg Val His ggt ggc gca gta gcc acc caa tct ttc caa acc aca gag ttg agc ttg 307 Gly Gly Ala Val Ala Thr Gln Ser Phe Gln Thr Thr Glu Leu Ser Leu gat act cgt ttc agg tct gca tcg tca gca aag tac tcc att gcc aag 355 Asp Thr Arg Phe Arg Ser Ala Ser Ser Ala Lys Tyr Ser Ile Ala Lys gca gcg atg cag ttc ctg ccc gct gag cat ggc gga ctg ttc ctc gat 403 Ala Ala Met Gln Phe Leu Pro Ala Glu His Gly Gly Leu Phe Leu Asp gcg gga act act gtt act gct ttg gcc gat ctc att tct gag cat cct Ala Gly Thr Thr Val Thr Ala Leu Ala Asp Leu Ile Ser Glu His Pro 105 110

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acc cgt cat Thr Arg His								L

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														att Ile		835
cag Gln	cag Gln	tgc Cys	acc Thr	cgt Arg 250	gag Glu	cag Gln	cgg Arg	gat Asp	ctt Leu 255	ttg Leu	cgt Arg	aat Asn	tcg Ser	cgc Arg 260	gcg Ala	883
														acc Thr		931
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<211> 282

<212> PRT

<213> Corynebacterium glutamicum

<400> 196

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Ala Glu Leu Phe Glu Val Ser Ala Met Thr Ile His Arg Asp Leu Glu 35 40 45

Ala Leu Ala Ala Asp Asn Leu Val Glu Arg Ile Arg Gly Gly Ala Arg 50 55 60

Ser Val Ser Pro Ser Met Ser Glu Leu Ala Val Glu Gln Arg Arg His 65 70 75 80

Leu His Arg Thr Val Lys Glu Ala Leu Cys Thr Ala Ala Ala Arg Leu  $85 \hspace{1cm} 90 \hspace{1cm} 95$ 

Ile Pro Glu Gly Ala Val Val Ala Ile Asp Asp Ser Thr Thr Leu Glu 100 105 110

Ser Leu Val Glu Lys Leu Pro Gln Arg Ser Pro Ser Ala Leu Ile Thr 115 120 125

His Ser Leu Lys Thr Met Ala Asp His Arg Val Arg Ala Gly Met Ser 130 135 140

Asp Ile Arg Leu Ile Ala Cys Ala Gly Leu Tyr Phe Ala Glu Thr Asp 145 150 155 160

Ser Phe Leu Gly Lys Ala Thr Ser Ala Gln Leu Asn Glu Leu Ser Ala 165 170 175

Asp Ile Ser Phe Val Ser Thr Thr Ala Val Arg Ala Thr Gly Glu Val 180 185 190

Pro Ala Leu Phe His Pro Asp Met Glu Ala Ala Asp Thr Lys Arg Ala 195 200 . 205

Leu Ile Gly Ile Gly Ser Val Arg Val Leu Val Val Asp Ser Ser Lys 210 215 220

Phe Gly Ser Ala Gly Val Phe Lys Val Ala Ser Ile Glu Glu Phe Asp 225 230 235 240

His Ile Ile Ile Asp Gln Gln Cys Thr Arg Glu Gln Arg Asp Leu Leu 245 250 255

Arg Asn Ser Arg Ala Gln Ile His Val Ile Asp His Asn Gly Asp Glu 260 265 270

Ile Leu Asp Thr Pro Thr Glu Glu Asp Phe 275 280

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284

185 190 180 ggc tac tgg tcc cgc gag acc ggt tat gtt cca gtt cgt aag gat gct 624 Gly Tyr Trp Ser Arg Glu Thr Gly Tyr Val Pro Val Arg Lys Asp Ala 195 200 gca tot gat oca gat cac gca gca tto oto gag gag aac oot gca tac 672 Ala Ser Asp Pro Asp His Ala Ala Phe Leu Glu Glu Asn Pro Ala Tyr 210 aac gtt gca gtg gag cag ctt cct gat acc cgt tcc cag gac aac ttc 720 Asn Val Ala Val Glu Gln Leu Pro Asp Thr Arg Ser Gln Asp Asn Phe 230 768 cgc gtg ctg ctg cca aac ggt gac cgc acc atc ggt gac gca ctg gag Arg Val Leu Leu Pro Asn Gly Asp Arg Thr Ile Gly Asp Ala Leu Glu aag atc tgc ctg act ggt gca gac atc gat gtc acc ctg gct gag gtt 816 Lys Ile Cys Leu Thr Gly Ala Asp Ile Asp Val Thr Leu Ala Glu Val gag acc aag ctg aac acc atc tac acc cgc gac atc gaa cca ctt att Glu Thr Lys Leu Asn Thr Ile Tyr Thr Arg Asp Ile Glu Pro Leu Ile 275 280 887 taatccgagc acttcagcta cac <210> 198 <211> 288 <212> PRT <213> Corynebacterium glutamicum <400> 198 Gly Gly His Tyr Gly Leu Pro Phe Ala Arg Ser Thr Val Leu Phe Tyr Tyr Asn Lys Asp Leu Trp Ala Lys Ala Gly Leu Glu Asp Arg Gly Pro Glu Ser Trp Glu Glu Phe Ser Glu Trp Gly Pro Lys Leu Gln Glu Ala Met Asp Ser Gly Phe Ala His Gly Trp Gly Asp Ala Thr Asn Tyr Leu Ser Trp Thr Phe Glu Gly Pro Met Trp Ser Leu Gly Gly Asn Tyr Ser Glu Gly Trp Glu Ser Arg Leu Thr Thr Pro Glu Thr Ile Arg Ala Val Glu Trp Leu Lys Ser Thr Val Asp Glu Gly Phe Ala Thr Val Ser Thr 100 Asp Val Thr Asn Glu Phe Ala Thr Gly Leu Ile Gly Ser Cys Ile Gln 120

Ser Thr Gly Asp Leu Ser Ser Val Ala Gly Ala Ala Ser Phe Asp Trp

135

130

Gly Val Ala Ala Leu Pro Asn Pro Thr Gly Glu Gly Ala Cys Pro Thr 160

Gly Gly Ala Gly Leu Gly Ile Pro Ser Gly Ile Ser Glu Gln Arg Gln Asp Asn Ala Leu Lys Phe Ile Asp Phe 185

Asp Asn Ala Leu Lys Phe Ile Asp Phe 185

Gly Tyr Trp Ser Arg Glu Thr 200 Tyr Val Pro Val Arg Lys Asp Ala 215

Ala Ser Asp Pro Asp His Ala Ala 215

Asn Val Ala Val Glu Gln 230

Asn Val Leu Leu Pro Asp Asn Gly Asp Arg Thr 250

Lys Ile Cys Leu Thr Gly Ala Asp Thr Ile Tyr Thr Arg Asp Ile Glu Pro Leu Ile 285

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							gca Ala									336
cag Gln	tcc Ser	acc Thr 115	ggt Gly	gat Asp	ctg Leu	tct Ser	tcg Ser 120	gtt Val	gcc Ala	ggc Gly	gct Ala	gca Ala 125	agc Ser	ttc Phe	gac Asp	384
tgg Trp	ggc Gly 130	gta Val	gca Ala	gca Ala	ctt Leu	cct Pro 135	aac Asn	cca Pro	acc Thr	ggc Gly	gag Glu 140	ggc Gly	gct Ala	tgc Cys	cca Pro	432
acc Thr 145	ggt Gly	ggc Gly	gca Ala	ggc Gly	ctg Leu 150	gga Gly	atc Ile	cca Pro	tct Ser	ggc Gly 155	atc Ile	tct Ser	gag Glu	cag Gln	cgt Arg 160	480
							atc Ile									528
act Thr	ggc Gly	tac Tyr	tgg Trp 180	tcc Ser	cgc Arg	gag Glu	acc Thr	ggt Gly 185	tat Tyr	gtt Val	cca Pro	gtt Val	cgt Arg 190	aag Lys	gat Asp	576
							gca Ala 200								gca Ala	624
tac Tyr	aac Asn 210	gtt Val	gca Ala	gtg Val	gag Glu	cag Gln 215	ctt Leu	cct Pro	gat Asp	acc Thr	cgt Arg 220	tcc Ser	cag Gln	gac Asp	aac Asn	672
							ggt Gly									720
gag Glu	aag Lys	atc Ile	tgc Cys	ctg Leu 245	act Thr	ggt Gly	gca Ala	gac Asp	atc Ile 250	gat Asp	gtc Val	acc Thr	ctg Leu	gct Ala 255	gag Glu	768
gtt Val	gag Glu	acc Thr	aag Lys 260	ctg Leu	aac Asn	acc Thr	atc Ile	tac Tyr 265	acc Thr	cgc Arg	gac Asp	atc Ile	gaa Glu 270	cca Pro	ctt Leu	816
att Ile	taa	teega	agc i	actto	caget	ta ca	ac									842

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<sup>&</sup>lt;211> 273

<sup>&</sup>lt;212> PRT

<sup>&</sup>lt;213> Corynebacterium glutamicum

<sup>&</sup>lt;400> 200

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Tyr Asn Val Ala Val Glu Gln Leu Pro Asp Thr Arg Ser Gln Asp Asn 210 215 220

Phe Arg Val Leu Leu Pro Asn Gly Asp Arg Thr Ile Gly Asp Ala Leu 225 230 235 240

Glu Lys Ile Cys Leu Thr Gly Ala Asp Ile Asp Val Thr Leu Ala Glu 245 250 255

Val Glu Thr Lys Leu Asn Thr Ile Tyr Thr Arg Asp Ile Glu Pro Leu 260 265 270

Ile

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cac																776
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Ala	Phe	Gln	Pro 20	Ala	Gly	Gly	Pro	Val 25	Lys	Pro	Trp	Asn	Lys 30	Pro	Asp	
Ala	Ser	Leu 35	Asn	Gln	Gln	Leu	Lys 40	Asn	Lys	Ser	Arg	Val 45	Arg	Thr	Gly	
Leu	Thr 50	Ile	Ala	Ile	Gly	Tyr 55	Val	Val	Val	Ile	Trp 60	Ala	Val	His	Leu	
Ala 65	Ser	Ile	Val	Ile	Ala 70	Leu	Leu	Thr	Gly	Phe 75	Asn	Leu	Thr	Asn	Phe 80	
Gly	Ile	His	Pro	Leu 85	Asp	Thr	Ser	Ala	Leu 90	Trp	Gly	Ile	Phe	Thr 95	Ser	
Pro	Leu	Leu	His 100	Gly	Ser	Phe	Ser	His 105	Leu	Ile	Gly	Asn	Thr 110	Val	Pro	
Gly	Phe	Ile 115	Phe	Ser	Phe	Leu	Ile 120	Gly	Met	Ser	Gly	Lys 125	Arg	Val	Phe	
Trp	Glu 130	Val	Thr	Ile	Ile	Ala 135	Gly	Leu	Ile	Gly	Gly 140	Leu	Gly	Thr	Trp	
Ile 145	Phe	Gly	Gly	Ile	Gly 150	Thr	Asn	His	Ile	Gly 155	Ala	Ser	Gly	Leu	Ile 160	
Tyr	Gly	Trp	Leu	Gly 165	Tyr	Leu	Ile	Val	Arg 170	Gly	Ile	Phe	Asn	Lys 175	Asp	
Ile	Lys	Gln	Phe 180	Leu	Leu	Gly	Ile	Val 185	Leu	Ala	Phe	Ile	Tyr 190	Ser	Gly	
Leu	Phe	Trp 195	Gly	Leu	Leu	Pro	Thr 200	Gln	Ile	Gly	Val	Ser 205	Trp	Gln	Gly	
His	Leu 210	Phe	Gly	Ala	Leu	Gly 215	Gly	Ile	Gly	Ala	Gly 220	Ala	Phe	Ile	Ala	
Ser 225	Asp	Asp	Pro	Ala	Ala 230	Leu	Lys	Ala	Lys	Lys 235	Gln	Gln	Lys	Lys	Leu 240	

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												aag Lys				691
												ggg Gly 210				739
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ctg																840
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His	Leu	Glu	Asn 20	Thr	Met	Thr	Ala	Phe 25	Gln	Ala	Ala	Ala	Pro 30	Ala	Asp	
Ala	Phe	Glu 35	Leu	Asp	Ile	His	Ala 40	Thr	Ala	Asp	Asn	Gln 45	Val	Val	Val	
Ile	His 50	Asp	Arg	Thr	Ala	Ala 55	Arg	Val	Ala	Ala	Pro 60	Asp	Ser	Leu	His	
Arg 65	Asp	Thr	Pro	Val	Ala 70	Arg	Leu	Ser	Ala	Ala 75	Gln	Ile	Lys	Glu	Ile 80	
Thr	Leu	Ile	Asp	Gly 85	Ser	Pro	Val	Pro	Thr 90	Leu	Glu	Glu	Val	Leu 95	Leu	
Gln	Thr	Ser	Leu 100	Pro	Ile	Gln	Val	Glu 105	Ile	Lys	Ser	Ala	Gly 110	Ala	Val	
Pro	Ala	Ala 115	Ala	Ala	Leu	Leu	Gln 120	Lys	Tyr	Pro	Glu	His 125	Leu	Glu	Arg	
Leu	Leu 130	Phe	Ile	Ser	Phe	Ile 135	Asp	Ala	Ala	Leu	Val 140	Glu	Ile	Val	Asp	•
Arg 145	Leu	Pro	Glu	Ala	Arg 150	Val	Gly	Ile	Leu	Arg 155	Asp	Ala	Ser	Met	Asp 160	
Asp	Leu	Arg	Ile	Leu	Asp	Tyr	Ile	Pro	Leu	Lys	Asn	Val	Gly	Ala	Ile	

165 170 175 Leu Pro Ser Trp Lys Ala Leu Asn Val Ala Ser Ile Ala Asp Leu His 185 Thr Lys Gly Ile Lys Val Gly Cys Trp Thr Ile Arg Asp Glu Asn Ala 200 Phe Gly Ile Ala Gln Gln Ala Gly Val Asp Tyr Ala Thr Val Ser Asp Pro Ser Arg Phe Leu Ala Pro Ser Pro Ala Gly Glu Leu His Trp 230 235 <210> 205 <211> 1314 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(1291) <223> RXA01436 <400> 205 gcctaaacaa accagtcaac gacctttccc gtggcgcaac agtccctgac atcgtcaaca 60 cagtagccat cacagcaatt caggcaggag gacgcagcta atg gca ttg gca ctt Met Ala Leu Ala Leu 1 gtt ttg aac tcc ggt tca tct tcc atc aaa ttc cag ctg gtc aac ccc 163 Val Leu Asn Ser Gly Ser Ser Ser Ile Lys Phe Gln Leu Val Asn Pro 10 15 gaa aac tot goo ato gao gag coa tat gtt tot ggt ott gtg gag cag 211 Glu Asn Ser Ala Ile Asp Glu Pro Tyr Val Ser Gly Leu Val Glu Gln 25 att ggt gag cca aac ggc cgc atc gta ctc aaa ata gag ggt gaa aaa Ile Gly Glu Pro Asn Gly Arg Ile Val Leu Lys Ile Glu Gly Glu Lys 40 tat acc cta gag aca ccc atc gca gat cac tcc gaa ggc cta aac ctg 307 Tyr Thr Leu Glu Thr Pro Ile Ala Asp His Ser Glu Gly Leu Asn Leu 55 gcg ttc gat ctc atg gac cag cac aac tgt ggt cct tcc caa ctg gaa 355 Ala Phe Asp Leu Met Asp Gln His Asn Cys Gly Pro Ser Gln Leu Glu 70 ate ace gea gtt gga cae ege gtg gte cae gge gga ate ttg tte tee 403 Ile Thr Ala Val Gly His Arg Val Val His Gly Gly Ile Leu Phe Ser 90 gca ccq qaa ctt atc act gat gaa atc gtg gaa atg atc cgc gat ctc 451

499

Ala Pro Glu Leu Ile Thr Asp Glu Ile Val Glu Met Ile Arg Asp Leu

att cca ctc gca cca ctg cac aac cct gca aac gtt gac ggc att gat

105

Ile Pr	o Leu 120	Ala	Pro	Leu	His	Asn 125	Pro	Ala	Asn	Val	Asp 130	Gly	Ile	Asp	
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acc gg Thr Gl 150															595
aac aa Asn Ly		_	-	-	-					-					643
ggc ac Gly Th			-				_	-			_		_	-	691
aag cc Lys Pr		Glu													739
gca tc Ala Se 21	r Met														787
ggt at Gly Me 230	_						-	_			_	-		-	835
att ga Ile As															883
atc ga Ile As															931
ctt tc Leu Se		Val		-		_	-	-		-	-		-		979
aat ga Asn As 29	p Gln														1027
cgc cg Arg Ar 310															1075
atc gt Ile Va															1123
gat go Asp Al	_	-		-	-	-						-			1171
cgt aa Arg As	-	_			_			-	_				_	-	1219

360 365 · 370

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<211> 397

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<213> Corynebacterium glutamicum

<400> 206

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Gly Leu Val Glu Gln Ile Gly Glu Pro Asn Gly Arg Ile Val Leu Lys 35 40 45

Ile Glu Gly Glu Lys Tyr Thr Leu Glu Thr Pro Ile Ala Asp His Ser
50 55 60

Glu Gly Leu Asn Leu Ala Phe Asp Leu Met Asp Gln His Asn Cys Gly 65 70 75 80

Pro Ser Gln Leu Glu Ile Thr Ala Val Gly His Arg Val Val His Gly 85 90 95

Gly Ile Leu Phe Ser Ala Pro Glu Leu Ile Thr Asp Glu Ile Val Glu 100 105 110

Met Ile Arg Asp Leu Ile Pro Leu Ala Pro Leu His Asn Pro Ala Asn 115 120 125

Val Asp Gly Ile Asp Val Ala Arg Lys Ile Leu Pro Asp Val Pro His 130 135 140

Val Ala Val Phe Asp Thr Gly Phe Phe His Ser Leu Pro Pro Ala Ala 145 150 155 160

Ala Leu Tyr Ala Ile Asn Lys Asp Val Ala Ala Glu His Gly İle Arg 165 170 175

Arg Tyr Gly Phe His Gly Thr Ser His Glu Phe Val Ser Lys Arg Val 180 185 190

Val Glu Ile Leu Glu Lys Pro Thr Glu Asp Ile Asn Thr Ile Thr Phe 195 200 205

His Leu Gly Asn Gly Ala Ser Met Ala Ala Val Gln Gly Gly Arg Ala 210 215 220

Val Asp Thr Ser Met Gly Met Thr Pro Leu Ala Gly Leu Val Met Gly 225 230 235 240

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Thr	Ala	Gly	Met 260	Ser	Ile	Asp	Glu	Ile 265	Asp	Asn	Leu	Leu	Asn 270	Lys	Lys	
Ser	Gly	Val 275	Lys	Gly	Leu	Ser	Gly 280	Val	Asn	Asp	Phe	Arg 285	Glu	Leu	Arg	
Glu	Met 290	Ile	Asp	Asn	Asn	Asp 295	Gln	Asp	Ala	Trp	Ser 300	Ala	Tyr	Asn	Ile	
Tyr 305	Ile	His	Gln	Leu	Arg 310	Arg	Tyr	Leu	Gly	Ser 315	Tyr	Met	Val	Ala	Leu 320	
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Gln	Phe	Val	Arg 340	Glu	Asp	Ala	Leu	Ala 345	Gly	Leu	Glu	Met	Tyr 350	Gly	Ile	
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gaa gct gta d Glu Ala Val 0 230												835
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att gaa cgc o Ile Glu Arg <i>l</i>	cgt gac cca cgc Arg Asp Pro Arg 25	gca gat gat gtg Ala Asp Asp Val 30	gtt att gat atc aa. Val Ile Asp Ile Ly: 35	a 211 s	
			atc cgc aac gaa tg Ile Arg Asn Glu Tr 50		
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			gaa tgc gaa cag tg Glu Cys Glu Gln Cy 100		
Val Ala Gly			aac gtc gga acc ta Asn Val Gly Thr Ty 115		
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			ggc atc acc acc ta Gly Ile Thr Thr Ty 16	r	
tcc cca atc o Ser Pro Ile o	gct cgc tgg aac Ala Arg Trp Asn 170	gtt aaa gaa ggc Val Lys Glu Gly 175	gac aaa gta gca gt Asp Lys Val Ala Va 180	c 643	
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Val Ile Asp Ile Lys Ala Ala Gly Ile Cys His Ser Asp Ile His Thr 35 40 45

Ile Arg Asn Glu Trp Gly Glu Ala His Phe Pro Leu Thr Val Gly His  $50 \hspace{1cm} 55 \hspace{1cm} 60$ 

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Lys Val Gly Asp Arg Val Gly Val Gly Cys Leu Val Asn Ser Cys Gly 85 90 95

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Asn Val Gly Thr Tyr Asn Ser Asn Asp Val Asp Gly Thr Ile Thr Gln 115 120 125

Gly Gly Tyr Ala Glu Lys Val Val Val Asn Glu Arg Phe Leu Cys Ser 130 135 140

Ile Pro Glu Glu Leu Asn Phe Asp Val Ala Ala Pro Leu Cys Ala 145 150 155 160

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cag cca ctg agc ttc ggt gcg ctc atc ggc ggc gga aaa gtc ctc acc Gln Pro Leu Ser Phe Gly Ala Leu Ile Gly Gly Gly Lys Val Leu Thr 90 95 100	403
gga tcc aac att ggc ggc atc cct gaa acc cag gaa atg ctc gac ttc Gly Ser Asn Ile Gly Gly Ile Pro Glu Thr Gln Glu Met Leu Asp Phe 105 110 115	451
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gct att cag ttt go Ala Ile Gln Phe Al 185			
cgc ggt tta gag cg Arg Gly Leu Glu An 200		Ala Arg Gln Leu	
tac atc gat agc as Tyr Ile Asp Ser As 215			
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gga gtt gat ggg gg Gly Val Asp Gly G 265			
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Ile Ala Gly Thr Ile Ala Glu Ile Gly Glu Asn Val Ser Arg Trp Thr 65 70 75 80

Val Gly Asp Arg Val Ala Ile Gly Trp Phe Gly Gly Asn Cys Gly Asp 85 90 95

Cys Ala Phe Cys Arg Ala Gly Asp Pro Val His Cys Arg Glu Arg Lys 100 105 110

Ile Pro Gly Val Ser Tyr Ala Gly Gly Trp Ala Gln Asn Ile Val Val 115 120 125

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Pro Ala Pro Met Gly Cys Ala Gly Val Thr Thr Phe Asn Ala Leu Arg 145 150 155 160

Asn Leu Lys Leu Asp Pro Gly Ala Ala Val Ala Val Phe Gly Ile Gly 165 170 175

Gly Leu Val Arg Leu Ala Ile Gln Phe Ala Ala Lys Met Gly Tyr Arg 180 185 190

Thr Ile Thr Ile Ala Arg Gly Leu Glu Arg Glu Glu Leu Ala Arg Gln 195 200 205

Leu Gly Ala Asn His Tyr Ile Asp Ser Asn Asp Leu His Pro Gly Gln 210 215 220

Ala Leu Phe Glu Leu Gly Gly Ala Asp Leu Ile Leu Ser Thr Ala Ser 225 230 235 240

Thr Thr Glu Pro Leu Ser Glu Leu Ser Thr Gly Leu Ser Ile Gly Gly 245 250 255

Gin Leu Thr Ile Ile Gly Val Asp Gly Gly Asp Ile Thr Val Ser Ala 260 265 270

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110

105

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	ggt Gly															691
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	tct Ser					_		-							_	835
	aag Lys	_			-	-	_		_	_				_		883
_	gag Glu				-								-	_	_	931
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	gcc Ala 295															1027
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	cgc Arg															1315
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His 145	Asn	Thr	Val	Ala	Tyr 150	His	Phe	Asn	Glu	Pro 155	Ile	Gly	Val	Val	Gly 160
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Leu	Gly	Val	Lys	Arg 325	Val	Gln	Asn	Ile	Lys 330	Leu	Gly	Asn	Pro	Leu 335	Asp
Thr	Glu	Thr	Met 340	Met	Gly	Ala	Gln	Ala 345	Ser	Gln	Glu	Gln	Met 350	Asp	Lys
Ile	Ser	Ser 355	Tyr	Leu	Lys	Ile	Gly 360	Pro	Glu	Glu	Gly	Ala 365	Gln	Thr	Leu
Thr	Gly 370	Gly	Lys	Val	Asn	Lys 375	Val	Asp	Gly	Met	Glu 380	Asn	Gly	Tyr	Tyr
Ile 385	Glu	Pro	Thr	Val	Phe 390	Arg	Gly	Thr	Asn	Asp 395	Met	Arg	Ile	Phe	Arg 400
Glu	Glu	Ile	Phe	Gly	Pro	Val	Leu	Ser	Val	Ala	Thr	Phe	Ser	Asp	Phe

410 405 415 Asp Glu Ala Ile Arg Ile Ala Asn Asp Thr Asn Tyr Gly Leu Gly Ala 425 Gly Val Trp Ser Arg Asp Gln Asn Thr Ile Tyr Arg Ala Gly Arg Ala 435 440 Ile Gln Ala Gly Arg Val Trp Val Asn Gln Tyr His Asn Tyr Pro Ala His Ser Ala Phe Gly Gly Tyr Lys Glu Ser Gly Ile Gly Arg Glu Asn 465 470 475 His Leu Met Met Leu Asn His Tyr Gln Gln Thr Lys Asn Leu Leu Val Ser Tyr Asp Pro Asn Pro Thr Gly Leu Phe 500 <210> 219 <211> 430 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(430) <223> RXN03061 <400> 219 ctgccaccac tggtcattgc agaggacact ctccgtgatg gtcttcaggt gttagtcgca 60 gccctagagc gcgaaaccgc gcaccagaag gtgggctaaa gtg tct ttg acc ttc Val Ser Leu Thr Phe 1 cca gta atc aac ccc agc gat ggc tcc acc atc acc gag cta gaa aac 163 Pro Val Ile Asn Pro Ser Asp Gly Ser Thr Ile Thr Glu Leu Glu Asn 10 cac gat tee ace cag tgg atg tee geg ete tet gat gea get 211 His Asp Ser Thr Gln Trp Met Ser Ala Leu Ser Asp Ala Val Ala Ala 25 ggt cct tca tgg gct gcg aaa act ccc cgc gaa aga tcc gtg gta ctc 259 Gly Pro Ser Trp Ala Ala Lys Thr Pro Arg Glu Arg Ser Val Val Leu 40 acc gca atc ttc gaa gca ctg acc gaa cgc gcc caa gaa ctt gca gag 307 Thr Ala Ile Phe Glu Ala Leu Thr Glu Arg Ala Gln Glu Leu Ala Glu 55 atc atc cac ctg gaa gct gga aaa tcc gat gca gaa gct ctt ggt gaa 355 Ile Ile His Leu Glu Ala Gly Lys Ser Asp Ala Glu Ala Leu Gly Glu 70 gtc gct tat ggt gca gaa tac ttc cgt tgg ttt gcg gaa gaa gca gtg 403 Val Ala Tyr Gly Ala Glu Tyr Phe Arg Trp Phe Ala Glu Glu Ala Val

430 cgc ctg ccc ggc cgc tac gga cag tca Arg Leu Pro Gly Arg Tyr Gly Gln Ser 105 <210> 220 <211> 110 <212> PRT <213> Corynebacterium glutamicum Val Ser Leu Thr Phe Pro Val Ile Asn Pro Ser Asp Gly Ser Thr Ile 10 Thr Glu Leu Glu Asn His Asp Ser Thr Gln Trp Met Ser Ala Leu Ser 30 25 Asp Ala Val Ala Ala Gly Pro Ser Trp Ala Ala Lys Thr Pro Arg Glu 40 Arg Ser Val Val Leu Thr Ala Ile Phe Glu Ala Leu Thr Glu Arg Ala 50 55 Gln Glu Leu Ala Glu Ile Ile His Leu Glu Ala Gly Lys Ser Asp Ala Glu Ala Leu Gly Glu Val Ala Tyr Gly Ala Glu Tyr Phe Arg Trp Phe Ala Glu Glu Ala Val Arg Leu Pro Gly Arg Tyr Gly Gln Ser 105 <210> 221 <211> 747 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(724) <223> RXN03150 <400> 221 tttaacagag tgcgtttcaa tgcctgtagt gttccggcaa ttttgaatgt cgttacggtt 60 acccaagget gaatteetga geteacettg tacaagatea gtg gaa gee eag tte Val Glu Ala Gln Phe acc tct ccc ctq ctc aac aat ggg caa acc tgt ttc ctt ggt acc cga 163 Thr Ser Pro Leu Leu Asn Asn Gly Gln Thr Cys Phe Leu Gly Thr Arg 15 atc ctt gct cca aaa tca cgt tac gcg gaa gta gtc gat gca ttc acc 211 Ile Leu Ala Pro Lys Ser Arg Tyr Ala Glu Val Val Asp Ala Phe Thr gct ttc gct ggc agc ctg cag gtt gga gtc acg tcc tcc cct gac act 259 Ala Phe Ala Gly Ser Leu Gln Val Gly Val Thr Ser Ser Pro Asp Thr

45 50 40 caq atc qqa ccq atq qcq act gcc cgg cag cgt gag cgc gtg gaa tcc 307 Gln Ile Gly Pro Met Ala Thr Ala Arg Gln Arg Glu Arg Val Glu Ser 60 tac att tee caa gge aaa aat get gga gee ege ate aet gte ggt gge 355 Tyr Ile Ser Gln Gly Lys Asn Ala Gly Ala Arg Ile Thr Val Gly Gly 70 75 80 age egt eea ega gat ett gae gee gga tte tte gtt gag eea aca gtg 403 Ser Arg Pro Arg Asp Leu Asp Ala Gly Phe Phe Val Glu Pro Thr Val 100 90 95 tto goo gat gta gad aat ogo goa goo att goo daa gat gaa atd tto 451 Phe Ala Asp Val Asp Asn Arg Ala Ala Ile Ala Gln Asp Glu Ile Phe 115 105 110 gga ccg gtg ccc tct gtt gtt tcc tac caa gac gat gaa cac gcc atc 499 Gly Pro Val Pro Ser Val Val Ser Tyr Gln Asp Asp Glu His Ala Ile 120 125 caa cta gcc aac gat tcc gaa ttc ggt ctc ggc gga act gtc tgg acg 547 Gln Leu Ala Asn Asp Ser Glu Phe Gly Leu Gly Gly Thr Val Trp Thr 135 140 age gat eee gag ege get gea ttg gee ege ega gtt eae aea gga Ser Asp Pro Glu Arg Gly Ala Ala Leu Ala Arg Arg Val His Thr Gly 160 150 155 acc att ggc atc aac cgc tat atc cct gat ccc gcc gca cca ttt gga Thr Ile Gly Ile Asn Arg Tyr Ile Pro Asp Pro Ala Ala Pro Phe Gly 170 175 ggt gtg aaa aac agt ggc ctt ggc aga gaa ctc ggc ccc gaa ggt ctt 691 Gly Val Lys Asn Ser Gly Leu Gly Arg Glu Leu Gly Pro Glu Gly Leu 185 190 get tee tae caa gaa aee caa aee att tat ete taateeaaae tgeacetata 744 Ala Ser Tyr Gln Glu Thr Gln Thr Ile Tyr Leu 205 200 747 tat <210> 222 <211> 208 <212> PRT <213> Corynebacterium glutamicum Val Glu Ala Gln Phe Thr Ser Pro Leu Leu Asn Asn Gly Gln Thr Cys Phe Leu Gly Thr Arg Ile Leu Ala Pro Lys Ser Arg Tyr Ala Glu Val 25 Val Asp Ala Phe Thr Ala Phe Ala Gly Ser Leu Gln Val Gly Val Thr

40

Ser Ser Pro Asp Thr Gln Ile Gly Pro Met Ala Thr Ala Arg Gln Arg

35

60

Glu Arg Val Glu Ser Tyr Ile Ser Gln Gly Lys Asn Ala Gly Ala Arg

55

Ile Thr Val Gly Gly Ser Arg Pro Arg Asp Leu Asp Ala Gly Phe Phe 85 90 95

Val Glu Pro Thr Val Phe Ala Asp Val Asp Asn Arg Ala Ala Ile Ala 100 105 110

Gln Asp Glu Ile Phe Gly Pro Val Pro Ser Val Val Sér Tyr Gln Asp 115 120 125

Asp Glu His Ala Ile Gln Leu Ala Asn Asp Ser Glu Phe Gly Leu Gly 130 135 140

Gly Thr Val Trp Thr Ser Asp Pro Glu Arg Gly Ala Ala Leu Ala Arg 145 150 155 160

Arg Val His Thr Gly Thr Ile Gly Ile Asn Arg Tyr Ile Pro Asp Pro 165 170 175

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Gly Pro Glu Gly Leu Ala Ser Tyr Gln Glu Thr Gln Thr Ile Tyr Leu 195 200 205

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1 5 10 15

cag ggc ttg gtc tca atc acc acc act cga gat gca gag cta tcg 96 Gln Gly Leu Val Ser Ile Ile Thr Thr Thr Arg Asp Ala Glu Leu Ser 20 25 30

gca gaa ctc atg gct gat cct cgc ttg gct aaa gtc acc ttc act gga 144 Ala Glu Leu Met Ala Asp Pro Arg Leu Ala Lys Val Thr Phe Thr Gly 35 40

tca acc aac gtg gga cgc atc ctg gtc cgc caa tcc gcg gac cga ctg 192 Ser Thr Asn Val Gly Arg Ile Leu Val Arg Gln Ser Ala Asp Arg Leu 50 55 60

ctg cgc acc tcc atg gaa ctc ggc gga aat gca gct ttt gtt atc gac 240

Leu 65	Arg	Thr	Ser	Met	Glu 70	Leu	Gly	Gly	Asn	Ala 75	Ala	Phe	Val	Ile	Asp 80	
					gac Asp											288
ctc Leu	cgc Arg	aac Asn	gcc Ala 100	ggc Gly	caa Gln	gta Val	tgc Cys	atc Ile 105	gca Ala	gct Ala	aac Asn	cgt Arg	ttc Phe 110	ttg Leu	gtt Val	336
					gcc Ala											384
					Gly Ggg											432
					gat Asp 150											480
					ccc Pro											528
	-	-	-	-	cct Pro	-	_							-		576
					acc Thr											624
					tcc Ser											672
					gca Ala 230											720
					aac Asn											768
					caa Gln											816
		-	-		ctc Leu			_			-	-	_			858
tga	cacat	cga q	gctgi	tacg	gt ga	aa										881

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<213> Corynebacterium glutamicum

<400> 224

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Gln Gly Leu Val Ser Ile Ile Thr Thr Thr Arg Asp Ala Glu Leu Ser 20 25 30

Ala Glu Leu Met Ala Asp Pro Arg Leu Ala Lys Val Thr Phe Thr Gly 35 40 45

Ser Thr Asn Val Gly Arg Ile Leu Val Arg Gln Ser Ala Asp Arg Leu 50 55 60

Leu Arg Thr Ser Met Glu Leu Gly Gly Asn Ala Ala Phe Val Ile Asp 65 70 75 80

Glu Ala Ala Asp Leu Asp Glu Ala Val Ser Gly Ala Ile Ala Ala Lys 85 90 95

Leu Arg Asn Ala Gly Gln Val Cys Ile Ala Ala Asn Arg Phe Leu Val 100 105 110

His Glu Ser Arg Ala Ala Glu Phe Thr Ser Lys Leu Ala Thr Ala Met 115 120 125

Gln Asn Thr Pro Ile Gly Pro Val Ile Ser Ala Arg Gln Arg Asp Arg 130 135 140

Ile Ala Ala Leu Val Asp Glu Ala Ile Thr Asp Gly Ala Arg Leu Ile 145 150 155 160

Ile Gly Gly Glu Val Pro Asp Gly Ser Gly Phe Phe Tyr Pro Ala Thr 165 170 175

Ile Leu Ala Asp Val Pro Ala Gln Ser Arg Ile Val His Glu Glu Ile 180 185 190

Phe Gly Pro Val Ala Thr Ile Ala Thr Phe Thr Asp Leu Ala Glu Gly
195 200 205

Val Ala Gln Ala Asn Ser Thr Glu Phe Gly Leu Ala Ala Tyr Gly Phe 210 215 220

Ser Asn Asn Val Lys Ala Thr Gln Tyr Met Ala Glu His Leu Glu Ala 225 230 235 240

Gly Met Val Gly Ile Asn Arg Gly Ala Ile Ser Asp Pro Ala Ala Pro 245 250 255

Phe Gly Gly Ile Gly Gln Ser Gly Phe Gly Arg Glu Gly Gly Thr Glu 260 265 270

Gly Ile Glu Glu Tyr Leu Ser Val Arg Tyr Leu Ala Leu Pro 275 280 285

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<212> DNA

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140 135 145 ace gae gaa aac tte gae eag act ttg eag gtt aac ete tat ggt agt 595 Thr Asp Glu Asn Phe Asp Gln Thr Leu Gln Val Asn Leu Tyr Gly Ser 150 155 ttt cgg gtt acc aaa gca gct ata cct cat ctg aag ccc gga tca tcg 643 Phe Arg Val Thr Lys Ala Ala Ile Pro His Leu Lys Pro Gly Ser Ser 170 ata atc ttt aca tcg tcc att cag gcg tac caa cct tcg gaa acc ctc 691 Ile Ile Phe Thr Ser Ser Ile Gln Ala Tyr Gln Pro Ser Glu Thr Leu 185 ttg gat tac gcc atg act aag gcg gca ttg aac aat ttg tca aag ggc 739 Leu Asp Tyr Ala Met Thr Lys Ala Ala Leu Asn Asn Leu Ser Lys Gly 200 205 ttg gca agt agt ctg ata ggc gat ggc att cgg gta aat tct gta gcc 787 Leu Ala Ser Ser Leu Ile Gly Asp Gly Ile Arg Val Asn Ser Val Ala 215 220 225 cca ggt cct ttc tgg acg ccg ttg caa ccc agc cat ggt cag cca caa 835 Pro Gly Pro Phe Trp Thr Pro Leu Gln Pro Ser His Gly Gln Pro Gln 235 230 gag aaa ata gaa gga ttt ggc cag cac gct ccg att gga aga gcg ggt 883 Glu Lys Ile Glu Gly Phe Gly Gln His Ala Pro Ile Gly Arg Ala Gly 250 255 260 cac cct gtt gag ttg gca ggt gcg tac gtt ttt ctc gct tct gac gaa 931 His Pro Val Glu Leu Ala Gly Ala Tyr Val Phe Leu Ala Ser Asp Glu 265 270 gcc agc tat gtg gta gga gaa acc ctg gga gtc aca ggt ggg acg ccc Ala Ser Tyr Val Val Gly Glu Thr Leu Gly Val Thr Gly Gly Thr Pro 280 285 acc cca tagtoggtac aagoggaato act 1008 Thr Pro 295 <210> 318 <211> 295 <212> PRT <213> Corynebacterium glutamicum <400> 318 Met Ile Ser Leu Leu Asn Asp Pro Arg Thr Leu Phe Pro Lys Val Asp Pro Pro Lys Gln Ser Gln Pro Glu Pro Gly Leu Asp Ile Lys Leu Ser 25 Pro Gln Ala Asp Ile Gly Leu Ser Ser Tyr Gln Gly Ser Gly Arg Leu 45 Lys Gly Arg Lys Ala Leu Ile Thr Gly Gly Asp Ser Gly Ile Gly Ala 55

Ala Val Ala Ile Ala Tyr Ala Arg Glu Gly Ala Asp Val Ala Ile Ala Tyr Leu Pro Glu Glu Gln Ala Asp Ala Asp Arg Val Leu Gln Ala Ile 90 Glu Glu Thr Gly Gln Lys Ala Phe Ser Phe Pro Gly Asp Leu Arg Asp Pro Glu Tyr Cys Arg Ser Leu Val Gln Glu Thr Val Asn Ala Leu Gly 120 Gly Leu Asp Ile Leu Val Asn Asn Ala Ser Arg Gln Val Trp Ala Pro 135 Gly Leu Thr Glu Ile Thr Asp Glu Asn Phe Asp Gln Thr Leu Gln Val 155 145 150 Asn Leu Tyr Gly Ser Phe Arg Val Thr Lys Ala Ala Ile Pro His Leu 170 Lys Pro Gly Ser Ser Ile Ile Phe Thr Ser Ser Ile Gln Ala Tyr Gln 180 Pro Ser Glu Thr Leu Leu Asp Tyr Ala Met Thr Lys Ala Ala Leu Asn 200 Asn Leu Ser Lys Gly Leu Ala Ser Ser Leu Ile Gly Asp Gly Ile Arg Val Asn Ser Val Ala Pro Gly Pro Phe Trp Thr Pro Leu Gln Pro Ser 235 His Gly Gln Pro Gln Glu Lys Ile Glu Gly Phe Gly Gln His Ala Pro Ile Gly Arg Ala Gly His Pro Val Glu Leu Ala Gly Ala Tyr Val Phe 265 Leu Ala Ser Asp Glu Ala Ser Tyr Val Val Gly Glu Thr Leu Gly Val 275 280 Thr Gly Gly Thr Pro Thr Pro 290 295 <210> 319 <211> 1605 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(1582) <223> RXN01049 <400> 319 aagcacaqca attgagcaat actcccatgc atgttttcgc gtgatcacgc tatatcctta 60 aagaatatto tttattagto agacotttaa aggaaacott atg gga toa att coa Met Gly Ser Ile Pro

aca atq tcc atc cct ttt qat qac tca cgt gga cct tat gtc ctt gct 163 Thr Met Ser Ile Pro Phe Asp Asp Ser Arg Gly Pro Tyr Val Leu Ala 10 atg gat att ggt tcc act gca tca cga ggt gga ctt tat gat gct tcc 211 Met Asp Ile Gly Ser Thr Ala Ser Arg Gly Gly Leu Tyr Asp Ala Ser ggc tgc cca atc aaa ggc acc aag cag cgc gaa tcc cat gaa ttc acc 259 Gly Cys Pro Ile Lys Gly Thr Lys Gln Arg Glu Ser His Glu Phe Thr acc ggt gag ggc gtt tcc acc att gat gct gac cag gtg gtt tcg gag 307 Thr Gly Glu Gly Val Ser Thr Ile Asp Ala Asp Gln Val Val Ser Glu 55 60 atc acc tca gtt att aat ggc att ttg aac gcg gct gat cat cac aac 355 Ile Thr Ser Val Ile Asn Gly Ile Leu Asn Ala Ala Asp His His Asn 80 70 403 atc aaa gat cag atc gcc gct gtc gcg cta gat tct ttt gca tcc tca Ile Lys Asp Gln Ile Ala Ala Val Ala Leu Asp Ser Phe Ala Ser Ser 95 100 90 tta atc ttg gtc gat ggt gaa ggc aat gcg ctc acc ccg tgc att acc 451 Leu Ile Leu Val Asp Gly Glu Gly Asn Ala Leu Thr Pro Cys Ile Thr 115 105 tac gcg gat tct cgt tct gca cag tat gtg gag cag ctg cgc gcg gaa 499 Tyr Ala Asp Ser Arg Ser Ala Gln Tyr Val Glu Gln Leu Arg Ala Glu 120 ate gat gag aag gee tae cae gge ege ace gge gte tge ttg cae ace Ile Asp Glu Lys Ala Tyr His Gly Arg Thr Gly Val Cys Leu His Thr 135 tee tae cae cea teg ege ttg etg tgg etg aaa aet gag tte gag aaa 595 Ser Tyr His Pro Ser Arg Leu Leu Trp Leu Lys Thr Glu Phe Glu Lys 150 155 gag tto aac aaa goo aag tat gtg atg acc atc ggt gag tac gtc tac 643 Glu Phe Asn Lys Ala Lys Tyr Val Met Thr Ile Gly Glu Tyr Val Tyr 170 ttc aaa ctt gca ggc atc acc gga atg gct act tcg att gcc gcg tgg 691 Phe Lys Leu Ala Gly Ile Thr Gly Met Ala Thr Ser Ile Ala Ala Trp 190 185 agt ggc att ttg gac gcc cat acc ggc gaa ctt gat ctg act atc ttg 739 Ser Gly Ile Leu Asp Ala His Thr Gly Glu Leu Asp Leu Thr Ile Leu 200 205 787 gag cac atc ggt gtt gat ccg gct ctg ttc ggt gag atc aga aac cct Glu His Ile Gly Val Asp Pro Ala Leu Phe Gly Glu Ile Arg Asn Pro 215 220 gat gaa cca gcc acc gat gcc aaa gtt gtc gac aaa aag tgg aag cac 835 Asp Glu Pro Ala Thr Asp Ala Lys Val Val Asp Lys Lys Trp Lys His 230

ctg gaa gaa ato Leu Glu Glu Ilo			e Pro Asp Gly		
aac att ggc cca Asn Ile Gly Pro 26	Gly Ala Val				
gct aca tcc ggg Ala Thr Ser Gly 280					
atc ccc tct ggd Ile Pro Ser Gly 295	c ctg tgg tgt / Leu Trp Cys 300	Tyr Arg Va	t tee ege gae 1 Ser Arg Asp 305	cag tgc at Gln Cys Il	c 1027 e
gtt ggt ggc gca Val Gly Gly Ala 310					u
cgc acc att ato Arg Thr Ile Ile			p Glu Val Leu		
ccc ctc gaa gge Pro Leu Glu Gl 34	Thr Pro Ala				
tcc atc ggc tgc Ser Ile Gly Trp 360					
gaa caa acc gge Glu Gln Thr Gly 375	-	Leu Trp Ar			
gca ctc tcc tac Ala Leu Ser Ty: 390					a
gcc cct gaa cgc Ala Pro Glu Arc	y Val Ile Ala	Ser Gly Ar	a gtc tcc acc g Val Ser Thr 5	Asp His Pr	a 1363 o
gaa ttc ctc gc Glu Phe Leu Ala 42	Met Leu Ser				
ctg gaa atg aad Leu Glu Met Ly: 440					
gag cag ctc gad Glu Gln Leu Glo 455		Thr Arg Al			
acg cat cag cc Thr His Gln Pro 470					u

PCT/IB00/00943 WO 01/00844

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1602

1605 cgc

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Ser His Glu Phe Thr Thr Gly Glu Gly Val Ser Thr Ile Asp Ala Asp

Gln Val Val Ser Glu Ile Thr Ser Val Ile Asn Gly Ile Leu Asn Ala 70

Ala Asp His His Asn Ile Lys Asp Gln Ile Ala Ala Val Ala Leu Asp 90

Ser Phe Ala Ser Ser Leu Ile Leu Val Asp Gly Glu Gly Asn Ala Leu

Thr Pro Cys Ile Thr Tyr Ala Asp Ser Arg Ser Ala Gln Tyr Val Glu

Gln Leu Arg Ala Glu Ile Asp Glu Lys Ala Tyr His Gly Arg Thr Gly

Val Cys Leu His Thr Ser Tyr His Pro Ser Arg Leu Leu Trp Leu Lys 155

Thr Glu Phe Glu Lys Glu Phe Asn Lys Ala Lys Tyr Val Met Thr Ile

Gly Glu Tyr Val Tyr Phe Lys Leu Ala Gly Ile Thr Gly Met Ala Thr

Ser Ile Ala Ala Trp Ser Gly Ile Leu Asp Ala His Thr Gly Glu Leu

Asp Leu Thr Ile Leu Glu His Ile Gly Val Asp Pro Ala Leu Phe Gly 215

Glu Ile Arg Asn Pro Asp Glu Pro Ala Thr Asp Ala Lys Val Val Asp 225

Lys Lys Trp Lys His Leu Glu Glu Ile Pro Trp Phe His Ala Ile Pro 250 245

Asp Gly Trp Pro Ser Asn Ile Gly Pro Gly Ala Val Asp Ser Lys Thr

Val Ala Val Ala Ala Ala Thr Ser Gly Ala Met Arg Val Ile Leu Pro Ser Val Pro Glu Gln Ile Pro Ser Gly Leu Trp Cys Tyr Arg Val Ser Arg Asp Gln Cys Ile Val Gly Gly Ala Leu Asn Asp Val Gly Arg Ala Val Thr Trp Leu Glu Arg Thr Ile Ile Lys Pro Glu Asn Leu Asp Glu Val Leu Ile Arg Glu Pro Leu Glu Gly Thr Pro Ala Val Leu Pro Phe Phe Ser Gly Glu Arg Ser Ile Gly Trp Ala Ala Ser Ala Gln Ala Thr Ile Thr Asn Ile Gln Glu Gln Thr Gly Pro Glu His Leu Trp Arg Gly Val Phe Glu Ala Leu Ala Leu Ser Tyr Gln Arg Val Trp Glu His Met 390 395 Gly Lys Ala Gly Ala Ala Pro Glu Arg Val Ile Ala Ser Gly Arg Val 405 410 415 Ser Thr Asp His Pro Glu Phe Leu Ala Met Leu Ser Asp Ala Leu Asp 420 425 Thr Pro Val Ile Pro Leu Glu Met Lys Arg Ala Thr Leu Arg Gly Thr 435 440 Ala Leu Ile Val Leu Glu Gln Leu Glu Pro Gly Gly Thr Arg Ala Thr 455 Pro Pro Phe Gly Thr Thr His Gln Pro Arg Phe Ala His His Tyr Ser Lys Ala Arg Glu Leu Phe Asp Ala Leu Tyr Leu Lys Leu Val 485 490 <210> 321 <211> 1134 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(1111) <223> FRXA01049 <400> 321 cacagtatgt ggagcagetg cgcgcggaaa tcgatgagaa ggcctaccac ggccgcaccg 60 gegtetgett geacacetee taccacecat egegettget gtg gtg aaa act gag Val Val Lys Thr Glu

	•	-, - , 0												
									1				5	
	gag Glu													163
	gtc Val													211
	gcg Ala													259
	atc Ile 55													307
	aac Asn													355
	aag Lys													403
	cct Pro													451
-	gcc Ala	_	-		 _	_	_				-	_	_	499
	gaa Glu 135	_			 _		-		-	_		_	-	547
	tgc Cys													595
	ctg Leu													643
	cgc Arg													691
	gaa Glu	_			 -	-			-	-	_			739
	att Ile 215	_	-			-		_		_		_		787
-	gcc Ala		_		_	_	-		_		-			835

gcc ggc gca Ala Gly Ala											883
gac cac cca Asp His Pro			Met I								931
gtc atc cct Val Ile Pro 280											979
atc gtc ctt Ile Val Leu 295			Pro C		Thr						1027
ttc ggc acg Phe Gly Thr 310											1075
aga gag ctt Arg Glu Leu							tago	tttt	cg		1121
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Arg Val Ser Arg Asp Gln Cys Ile Val Gly Gly Ala Leu Asn Asp Val

Gly Arg Ala Val Thr Trp Leu Glu Arg Thr Ile Ile Lys Pro Glu Asn Leu Asp Glu Val Leu Ile Arg Glu Pro Leu Glu Gly Thr Pro Ala Val Leu Pro Phe Phe Ser Gly Glu Arg Ser Ile Gly Trp Ala Ala Ser Ala Gln Ala Thr Ile Thr Asn Ile Gln Glu Gln Thr Gly Pro Glu His Leu Trp Arg Gly Val Phe Glu Ala Leu Ala Leu Ser Tyr Gln Arg Val Trp 235 Glu His Met Gly Lys Ala Gly Ala Ala Pro Glu Arg Val Ile Ala Ser Gly Arg Val Ser Thr Asp His Pro Glu Phe Leu Ala Met Leu Ser Asp 265 Ala Leu Asp Thr Pro Val Ile Pro Leu Glu Met Lys Arg Ala Thr Leu 280 Arg Gly Thr Ala Leu Ile Val Leu Glu Gln Leu Glu Pro Gly Gly Thr 295 Arg Ala Thr Pro Pro Phe Gly Thr Thr His Gln Pro Arg Phe Ala His His Tyr Ser Lys Ala Arg Glu Leu Phe Asp Ala Leu Tyr Leu Lys Leu Val <210> 323 <211> 597 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(574) <223> FRXA01050 <400> 323 aagcacagca attgagcaat actcccatgc atgttttcgc gtgatcacgc tatatcctta 60 115 aaqaatatto tttattagto agacotttaa aggaaacott atg gga toa att coa Met Gly Ser Ile Pro 163 aca atg tee ate eet ttt gat gae tea egt gga eet tat gte ett get Thr Met Ser Ile Pro Phe Asp Asp Ser Arg Gly Pro Tyr Val Leu Ala 10 15

								cga Arg 30								211	
								cag Gln								259	
acc Thr	ggt Gly 55	gag Glu	ggc Gly	gtt Val	tcc Ser	acc Thr 60	att Ile	gat Asp	gct Ala	gac Asp	cag Gln 65	gtg Val	gtt Val	tcg Ser	gag Glu	307	
								ttg Leu								355	
								gcg Ala								403	
								aat Asn 110								451	٠
								tat Tyr								499	
								cgc Arg								547	
				_	-	-	ctg Leu	tgg Trp	tgaa	aaact	tga q	gttc	gagaa	aa		594	
gag																597	
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1				5					10					15			
Pro	Tyr	Val	Leu 20	Ala	Met	Asp	Ile	Gly 25	Ser	Thr	Ala	Ser	Arg 30	Gly	Gly		
Leu	Tyr	Asp 35	Ala	Ser	Gly	Cys	Pro 40	Ile	Lys	Gly	Thr	Lys 45	Gln	Arg	Glu		
Ser	His 50	Glu	Phe	Thr	Thr	Gly 55	Glu	Gly	Val	Ser	Thr 60	Ile	Asp	Ala	Asp		
Gln 65	Val	Val	Ser	Glu	Ile 70	Thr	Ser	Val	Ile	Asn 75	Gly	Ile	Leu	Asn	Ala 80		
Ala	Asp	His	His	Asn	Ile	Lys	Asp	Gln	Ile	Ala	Ala	Val	Ala	Leu	Asp		

85 90 95 Ser Phe Ala Ser Ser Leu Ile Leu Val Asp Gly Glu Gly Asn Ala Leu 105 Thr Pro Cys Ile Thr Tyr Ala Asp Ser Arg Ser Ala Gln Tyr Val Glu 120 Gln Leu Arg Ala Glu Ile Asp Glu Lys Ala Tyr His Gly Arg Thr Gly Val Cys Leu His Thr Ser Tyr His Pro Ser Arg Leu Leu Trp 150 <210> 325 <211> 1065 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(1042) <223> RXA00202 <400> 325 ctggcagcag attgtcatcg gttgtgtcat cgcgcttgcg gtgggcttcg atgtcatccg 60 aaacaaaacc tctaagtaat tcctgaaagg aaattttcac atg tac gct cgt aaa 115 Met Tyr Ala Arg Lys 1 ctt att gct ctg tcc gct tct gtc gtt ttg gct ttc agc ttg tct gct 163 Leu Ile Ala Leu Ser Ala Ser Val Val Leu Ala Phe Ser Leu Ser Ala 10 tgc aac cgt gaa tct tct ggc acc agc gca gac ggc ggt tct gcg gat 211 Cys Asn Arg Glu Ser Ser Gly Thr Ser Ala Asp Gly Gly Ser Ala Asp 25 ggg tcg atc acc ttg gct ctg tct acc cag acc aac ccg ttc ttt gtg 259 Gly Ser Ile Thr Leu Ala Leu Ser Thr Gln Thr Asn Pro Phe Phe Val 40 45 307 cag ctt cgt gat ggt gcc cag gaa aag gct gat gaa ttg ggc gtg acc Gln Leu Arg Asp Gly Ala Gln Glu Lys Ala Asp Glu Leu Gly Val Thr 55 ctc aat gtt cag gat gct tcc gat gac gct gca acg cag gcc aac cag 355 Leu Asn Val Gln Asp Ala Ser Asp Asp Ala Ala Thr Gln Ala Asn Gln 70 ctc aac aac gct gtc acc acc ggt gct ggc gtg gtg att gtc aac cca 403 Leu Asn Asn Ala Val Thr Thr Gly Ala Gly Val Val Ile Val Asn Pro 90 act gat tot gat gct gtg gtg ccg tcg gtg gaa gct ctc aac cag gct 451 Thr Asp Ser Asp Ala Val Val Pro Ser Val Glu Ala Leu Asn Gln Ala

gac att cct gtt gtg gct gtc gac cgt tcc tcc aat ggt ggc gag gtg

115

105

Asp Ile Pro 1	Val Val	Ala Val	. Asp 125	Arg	Ser	Ser	Asn	Gly 130	Gly	Glu	Val	
gcg tcc ttc of Ala Ser Phe 1 135			Asn									547
gca gcc ctg o Ala Ala Leu i 150												595
caa ggc att o Gln Gly Ile i												643
gaa gag gag a Glu Glu Glu :												691
acc gcc aac f Thr Ala Asn 1 200												739
ctg cag gca o Leu Gln Ala 1 215			Lys									787
atg gcg ttg ( Met Ala Leu ( 230												835
gtc atc gtt o												883
gaa gat gga G Glu Asp Gly i												931
gga gca aag ( Gly Ala Lys 7 280												979
gct gaa aca ( Ala Glu Thr ( 295			. Glu									1027
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•	3				10							

Phe Ser Leu Ser Ala Cys Asn Arg Glu Ser Ser Gly Thr Ser Ala Asp Gly Gly Ser Ala Asp Gly Ser Ile Thr Leu Ala Leu Ser Thr Gln Thr Asn Pro Phe Phe Val Gln Leu Arg Asp Gly Ala Gln Glu Lys Ala Asp Glu Leu Gly Val Thr Leu Asn Val Gln Asp Ala Ser Asp Asp Ala Ala Thr Gln Ala Asn Gln Leu Asn Asn Ala Val Thr Thr Gly Ala Gly Val Val Ile Val Asn Pro Thr Asp Ser Asp Ala Val Val Pro Ser Val Glu Ala Leu Asn Gln Ala Asp Ile Pro Val Val Ala Val Asp Arg Ser Ser Asn Gly Gly Glu Val Ala Ser Phe Val Ala Ser Asp Asn Val Ala Gly 135 Gly Ala Gln Ala Ala Ala Leu Ala Glu Ala Ile Gly Gly Glu Gly Glu Ile Leu Met Leu Gln Gly Ile Ala Gly Ser Ser Ala Ser Arg Asp 170 Arg Gly Gln Gly Phe Glu Glu Glu Ile Ala Lys His Glu Gly Ile Ser 185 Ile Val Ala Lys Gln Thr Ala Asn Phe Asp Arg Gly Glu Gly Leu Asp 195 200 Val Ala Thr Asn Leu Leu Gln Ala His Pro Asn Val Lys Ala Ile Phe Ala Glu Asn Asp Glu Met Ala Leu Gly Ala Ile Glu Ala Leu Gly Ala Arg Ala Gly Glu Asp Val Ile Val Val Gly Phe Asp Gly Thr Asn Asp Gly Leu Ala Ala Val Glu Asp Gly Arg Met Leu Ala Thr Val Ala Gln Gln Pro Glu Glu Leu Gly Ala Lys Ala Val Glu Glu Ala Ala Lys Leu 280 Leu Arg Gly Glu Asp Ala Glu Thr Glu Val Pro Val Glu Val Val Thr 300 Val Lys Leu Asp Asn Val Ala Asp Phe Lys 310

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gaa gag gcc acc aag cag gcg gaa tgg gcg ttg cag cat tcc acg gtg Glu Glu Ala Thr Lys Gln Ala Glu Trp Ala Leu Gln His Ser Thr Val 185 190 195

age ege gge aca cae ege gag ate ttg act act egt geg aac egt ege

Ser Arg Gly Thr His Arg Glu Ile Leu Thr Thr Arg Ala Asn Arg Arg

cac acc atc ttt gat ctg gac tac cga cca atg ttc tgg gaa tcc cca

His Thr Ile Phe Asp Leu Asp Tyr Arg Pro Met Phe Trp Glu Ser Pro

155

170

150

595

643

691

180